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**WO 02/07678 A2**

(54) Title: **MU-CONOPEPTIDES**

(57) Abstract: The present invention is to  $\mu$ -conopeptides, derivatives or pharmaceutically acceptable salts thereof. The present invention is further directed to the use of this peptide, derivatives thereof and pharmaceutically acceptable salts thereof for the treatment of disorders associated with voltage-gated sodium channels. Thus, the  $\mu$ -conopeptides or derivatives are useful as neuromuscular blocking agents, local anesthetic agents, analgesic agents and neuroprotective agents. The  $\mu$ -conopeptides are also useful for treating neuromuscular disorders. The invention is further directed to nucleic acid sequences encoding the  $\mu$ -conopeptides and encoding propeptides, as well as the propeptides.

TITLE OF THE INVENTION

MU-CONOPEPTIDES

CROSS-REFERENCE TO RELATED APPLICATIONS

5 [0001] The present application claims benefit under 35 USC §119(e) to U.S. provisional patent applications Serial No. 60/219,619 filed on 21 July 2000, Serial No. 60/245,157 filed on 3 November 2000, Serial No. 60/264,319 filed on 29 January 2001 and Serial No. 60/277,270 filed on 21 March 2001. Each of these applications is incorporated herein by reference.

10 [0002] This invention was made with Government support under Grant No. PO1 GM48677 awarded by the National Institute of General Medical Sciences, National Institutes of Health, Bethesda, Maryland. The United States Government has certain rights in the invention.

BACKGROUND OF THE INVENTION

15 [0003] The present invention is to  $\mu$ -conopeptides, derivatives or pharmaceutically acceptable salts thereof. The present invention is further directed to the use of this peptide, derivatives thereof and pharmaceutically acceptable salts thereof for the treatment of disorders associated with voltage-gated sodium channels. Thus, the  $\mu$ -conopeptides or derivatives are useful as neuromuscular blocking agents, local anesthetic agents, analgesic agents and 20 neuroprotective agents. The  $\mu$ -conopeptides are also useful for treating neuromuscular disorders. The invention is further directed to nucleic acid sequences encoding the  $\mu$ -conopeptides and encoding propeptides, as well as the propeptides.

25 [0004] The publications and other materials used herein to illuminate the background of the invention, and in particular, cases to provide additional details respecting the practice, are incorporated by reference, and for convenience are referenced in the following text by author and date and are listed alphabetically by author in the appended bibliography.

30 [0005] *Conus* is a genus of predatory marine gastropods (snails) which envenomate their prey. Venomous cone snails use a highly developed projectile apparatus to deliver their cocktail of toxic conotoxins into their prey. In fish-eating species such as *Conus magus* the cone detects the presence of the fish using chemosensors in its siphon and when close enough extends its proboscis and fires a hollow harpoon-like tooth containing venom into the fish. This immobilizes the fish and enables the cone snail to wind it into its mouth via an attached filament. For general information on *Conus* and their venom see the website address

http://grimwade.biochem.unimelb.edu.au/cone/referenc.html. Prey capture is accomplished through a sophisticated arsenal of peptides which target specific ion channel and receptor subtypes. Each *Conus* species venom appears to contain a unique set of 50-200 peptides. The composition of the venom differs greatly between species and between individual snails within each species, each optimally evolved to paralyse it's prey. The active components of the venom are small peptides toxins, typically 10-40 amino acid residues in length and are typically highly constrained peptides due to their high density of disulphide bonds.

[0006] The venoms consist of a large number of different peptide components that when separated exhibit a range of biological activities: when injected into mice they elicit a range of physiological responses from shaking to depression. The paralytic components of the venom that have been the focus of recent investigation are the  $\alpha$ -,  $\omega$ - and  $\mu$ -conotoxins. All of these conotoxins act by preventing neuronal communication, but each targets a different aspect of the process to achieve this. The  $\alpha$ -conotoxins target nicotinic ligand gated channels, the  $\mu$ -conotoxins target the voltage-gated sodium channels and the  $\omega$ -conotoxins target the voltage-gated calcium channels (Olivera et al., 1985; Olivera et al., 1990). For example a linkage has been established between  $\alpha$ -,  $\alpha$ A- &  $\phi$ -conotoxins and the nicotinic ligand-gated ion channel;  $\omega$ -conotoxins and the voltage-gated calcium channel;  $\mu$ -conotoxins and the voltage-gated sodium channel;  $\delta$ -conotoxins and the voltage-gated sodium channel;  $\kappa$ -conotoxins and the voltage-gated potassium channel; conantokins and the ligand-gated glutamate (NMDA) channel.

[0007] However, the structure and function of only a small minority of these peptides have been determined to date. For peptides where function has been determined, three classes of targets have been elucidated: voltage-gated ion channels; ligand-gated ion channels, and G-protein-linked receptors.

[0008] *Conus* peptides which target voltage-gated ion channels include those that delay the inactivation of sodium channels, as well as blockers specific for sodium channels, calcium channels and potassium channels. Peptides that target ligand-gated ion channels include antagonists of NMDA and serotonin receptors, as well as competitive and noncompetitive nicotinic receptor antagonists. Peptides which act on G-protein receptors include neuropeptides and vasopressin receptor agonists. The unprecedented pharmaceutical selectivity of conotoxins is at least in part defined by a specific disulfide bond frameworks combined with hypervariable amino acids within disulfide loops (for a review see McIntosh et al., 1998).

[0009] There are drugs used in the treatment of pain, which are known in the literature and to the skilled artisan. See, for example, Merck Manual, 16th Ed. (1992). However, there is a demand for more active analgesic agents with diminished side effects and toxicity and which are non-addictive. The ideal analgesic would reduce the awareness of pain, produce analgesia over a wide range of pain types, act satisfactorily whether given orally or parenterally, produce minimal or no side effects, be free from tendency to produce tolerance and drug dependence.

[0010] Due to the high potency and exquisite selectivity of the conopeptides, several are in various stages of clinical development for treatment of human disorders. For example, two *Conus* peptides are being developed for the treatment of pain. The most advanced is  $\omega$ -conotoxin MVIIA (ziconotide), an N-type calcium channel blocker (see Heading, C., 1999; U.S. Patent No. 5,859,186).  $\omega$ -Conotoxin MVIIA, isolated from *Conus magus*, is approximately 1000 times more potent than morphine, yet does not produce the tolerance or addictive properties of opiates.  $\omega$ -Conotoxin MVIIA has completed Phase III (final stages) of human clinical trials and has been approved as a therapeutic agent.  $\omega$ -Conotoxin MVIIA is introduced into human patients by means of an implantable, programmable pump with a catheter threaded into the intrathecal space. Preclinical testing for use in post-surgical pain is being carried out on another *Conus* peptide, contulakin-G, isolated from *Conus geographus* (Craig et al. 1999). Contulakin-G is a 16 amino acid O-linked glycopeptide whose C-terminus resembles neuropeptides. It is an agonist of neuropeptides receptors, but appears significantly more potent than neuropeptides in inhibiting pain in *in vivo* assays.

[0011] In view of a large number of biologically active substances in *Conus* species it is desirable to further characterize them and to identify peptides capable of treating disorders involving voltage gated ion channels, such as stroke and pain. Surprisingly, and in accordance with this invention, Applicants have discovered novel conotoxins that can be useful for the treatment of disorders involving voltage gated ion channels and could address a long felt need for a safe and effective treatment.

#### SUMMARY OF THE INVENTION

[0012] The present invention is to  $\mu$ -conopeptides, derivatives or pharmaceutically acceptable salts thereof. The present invention is further directed to the use of this peptide, derivatives thereof and pharmaceutically acceptable salts thereof for the treatment of disorders associated with voltage-gated sodium channels. Thus, the  $\mu$ -conopeptides or derivatives are

useful as neuromuscular blocking agents, local anesthetic agents, analgesic agents and neuroprotective agents. The  $\mu$ -conopeptides are also useful for treating neuromuscular disorders. The invention is further directed to nucleic acid sequences encoding the  $\mu$ -conopeptides and encoding propeptides, as well as the propeptides.

5 [0013] More specifically, the present invention is directed to  $\mu$ -conopeptides, having the amino acid sequences set forth in Tables 1 and 2 below.

[0014] The present invention is also directed to derivatives or pharmaceutically acceptable salts of the  $\mu$ -conopeptides or the derivatives. Examples of derivatives include peptides in which the Arg residues may be substituted by Lys, ornithine, homoarginine, nor-Lys, 10 N-methyl-Lys, N,N-dimethyl-Lys, N,N,N-trimethyl-Lys or any synthetic basic amino acid; the Lys residues may be substituted by Arg, ornithine, homoarginine, nor-Lys, or any synthetic basic amino acid; the Tyr residues may be substituted with meta-Tyr, ortho-Tyr, nor-Tyr, mono-halo-Tyr, di-halo-Tyr, O-sulpho-Tyr, O-phospho-Tyr, nitro-Tyr or any synthetic hydroxy containing amino acid; the Ser residues may be substituted with Thr or any synthetic hydroxylated amino acid; the Thr residues may be substituted with Ser or any synthetic hydroxylated amino acid; the Phe residues may be substituted with any synthetic aromatic amino acid; the Trp residues may be substituted with Trp (D), neo-Trp, halo-Trp (D or L) or any aromatic synthetic amino acid; and the Asn, Ser, Thr or Hyp residues may be glycosylated. The halogen may be iodo, chloro, fluoro or bromo; preferably iodo for halogen substituted-Tyr and bromo for halogen-substituted 15 Trp. The Tyr residues may also be substituted with the 3-hydroxyl or 2-hydroxyl isomers (meta-Tyr or ortho-Tyr, respectively) and corresponding O-sulpho- and O-phospho-derivatives. The acidic amino acid residues may be substituted with any synthetic acidic amino acid, e.g., tetrazolyl derivatives of Gly and Ala. The aliphatic amino acids may be substituted by synthetic derivatives bearing non-natural aliphatic branched or linear side chains  $C_nH_{2n+2}$  up to and 20 including n=8. The Met residues may be substituted by norleucine (Nle). The Cys residues may be in D or L configuration and may optionally be substituted with homocysteine (D or L).

[0015] Examples of synthetic aromatic amino acid include, but are not limited to, nitro-Phe, 4-substituted-Phe wherein the substituent is  $C_1-C_3$ , alkyl, carboxyl, hydroxymethyl, sulphomethyl, halo, phenyl, -CHO, -CN, -SO<sub>3</sub>H and -NHA. Examples of synthetic hydroxy 25 containing amino acid, include, but are not limited to, such as 4-hydroxymethyl-Phe, 4-hydroxyphenyl-Gly, 2,6-dimethyl-Tyr and 5-amino-Tyr. Examples of synthetic basic amino acids include, but are not limited to, N-1-(2-pyrazolinyl)-Arg, 2-(4-piperinyl)-Gly, 2-(4-

piperinyl)-Ala, 2-[3-(2S)pyrrolininy]-Gly and 2-[3-(2S)pyrrolininy]-Ala. These and other synthetic basic amino acids, synthetic hydroxy containing amino acids or synthetic aromatic amino acids are described in Building Block Index, Version 3.0 (1999 Catalog, pages 4-47 for hydroxy containing amino acids and aromatic amino acids and pages 66-87 for basic amino acids; see also <http://www.amino-acids.com>), incorporated herein by reference, by and available from RSP Amino Acid Analogues, Inc., Worcester, MA. Examples of synthetic acid amino acids include those derivatives bearing acidic functionality, including carboxyl, phosphate, sulfonate and synthetic tetrazolyl derivatives such as described by Ornstein et al. (1993) and in U.S. Patent No. 5,331,001, each incorporated herein by reference.

[0016] Optionally, in the  $\mu$ -conopeptides of the present invention, the Asn residues may be modified to contain an N-glycan and the Ser, Thr and Hyp residues may be modified to contain an O-glycan (e.g., g-N, g-S, g-T and g-Hyp). In accordance with the present invention, a glycan shall mean any N-, S- or O-linked mono-, di-, tri-, poly- or oligosaccharide that can be attached to any hydroxy, amino or thiol group of natural or modified amino acids by synthetic or enzymatic methodologies known in the art. The monosaccharides making up the glycan can include D-allose, D-altrose, D-glucose, D-mannose, D-gulose, D-idose, D-galactose, D-talose, D-galactosamine, D-glucosamine, D-N-acetyl-glucosamine (GlcNAc), D-N-acetyl-galactosamine (GalNAc), D-fucose or D-arabinose. These saccharides may be structurally modified, e.g., with one or more O-sulfate, O-phosphate, O-acetyl or acidic groups, such as sialic acid, including combinations thereof. The glycan may also include similar polyhydroxy groups, such as D-penicillamine 2,5 and halogenated derivatives thereof or polypropylene glycol derivatives. The glycosidic linkage is beta and 1-4 or 1-3, preferably 1-3. The linkage between the glycan and the amino acid may be alpha or beta, preferably alpha and is 1-.

[0017] Core O-glycans have been described by Van de Steen et al. (1998), incorporated herein by reference. Mucin type O-linked oligosaccharides are attached to Ser or Thr (or other hydroxylated residues of the present peptides) by a GalNAc residue. The monosaccharide building blocks and the linkage attached to this first GalNAc residue define the "core glycans," of which eight have been identified. The type of glycosidic linkage (orientation and connectivities) are defined for each core glycan. Suitable glycans and glycan analogs are described further in U.S. Serial No. 09/420,797 filed 19 October 1999 and in PCT Application No. PCT/US99/24380 filed 19 October 1999 (PCT Published Application No. WO 00/23092), each incorporated herein by reference. A preferred glycan is Gal( $\beta$ 1 $\rightarrow$ 3)GalNAc( $\alpha$ 1 $\rightarrow$ ).

[0018] Optionally, in the  $\mu$ -conopeptides described above, pairs of Cys residues may be replaced pairwise with isoteric lactam or ester-thioether replacements, such as Ser/(Glu or Asp), Lys/(Glu or Asp), Cys/(Glu or Asp) or Cys/Ala combinations. Sequential coupling by known methods (Barnay et al., 2000; Hruby et al., 1994; Bitan et al., 1997) allows replacement of native Cys 5 bridges with lactam bridges. Thioether analogs may be readily synthesized using halo-Ala residues commercially available from RSP Amino Acid Analogues.

[0019] The present invention is further directed to derivatives of the above peptides and peptide derivatives which are acyclic permutations in which the cyclic permutants retain the native bridging pattern of native toxin. See, for example, Craik et al. (2001).

[0020] The present invention is further directed to a method of treating disorders associated with voltage gated ion channel disorders in a subject comprising administering to the subject an effective amount of the pharmaceutical composition comprising a therapeutically effective amount of a  $\mu$ -conopeptide described herein or a pharmaceutically acceptable salt or solvate thereof. The present invention is also directed to a pharmaceutical composition comprising a therapeutically effective amount of a  $\mu$ -conopeptide described herein or a pharmaceutically acceptable salt or solvate thereof and a pharmaceutically acceptable carrier.

[0021] More specifically, the present invention is further directed to uses of these peptides or nucleic acids as described herein as neuromuscular blocking agents, local anesthetic agents, analgesic agents and neuroprotective agents. The  $\mu$ -conopeptides are also useful for 20 treating neuromuscular disorders.

[0022] The present invention is directed to the use of  $\mu$ -conopeptides as a local anesthetic for treating pain. The  $\mu$ -conopeptides have long lasting anesthetic activity and are particularly useful for spinal anesthesia, either administered acutely for post-operative pain or via an intrathecal pump for severe chronic pain situations. The  $\mu$ -conopeptides are also useful as 25 analgesics in chronic and neuropathic pain states, such as trigeminal neuralgia, diabetic neuropathy, post-herpetic neuralgia, neuroma pain and phantom limb pain. The  $\mu$ -conopeptides are also useful for treating burn pain and as ocular anesthetics.

[0023] The present invention is directed to the use of  $\mu$ -conopeptides as neuroprotectants. The  $\mu$ -conopeptides are useful for the treatment and alleviation of epilepsy 30 and as a general anticonvulsant agent. The  $\mu$ -conopeptides are also useful for treating neurodegenerative diseases, such as Amyotrophic Lateral Sclerosis (ALS). The  $\mu$ -conopeptides are further useful as cerebroprotectants, such as for reducing neurotoxic injury associated with

conditions of hypoxia, anoxia or ischemia which typically follows stroke, cerebrovascular accident, brain or spinal cord trauma, myocardial infarct, physical trauma, drowning, suffocation, perinatal asphyxia, or hypoglycemic events.

[0024] The present invention is directed to the use of  $\mu$ -conopeptides as neuromuscular blockers and for treating neuromuscular disorders. As such, the  $\mu$ -conopeptides are useful for providing relaxation of muscle, for treating benign essential blepharospasm and other forms of focal dystonia and for anti-wrinkle use.

[0025] More specifically, the present invention is also directed to nucleic acids which encode  $\mu$ -conopeptides of the present invention or which encodes precursor peptides for these  $\mu$ -conopeptides, as well as the precursor peptide. The nucleic acid sequences encoding the precursor peptides of other  $\mu$ -conopeptides of the present invention are set forth in Table 1. Table 1 also sets forth the amino acid sequences of these precursor peptides.

[0026] The present invention is further directed to the use of selectively radioiodinated or radiotriitated  $\mu$ -conopeptides for characterizing pore occlusion sites on different sodium channel subtypes or for use in screening assays.

[0027] The present invention is also directed to the use of  $\mu$ -conopeptides for screening small molecule libraries to identify small molecules that are selective blocking agents at specific sodium channel subtypes expressed in mammalian systems. In one embodiment, the blocking activity of a small molecule at a particular sodium channel subtype is compared to the blocking activity of a  $\mu$ -conopeptide at the same sodium channel subtype. In a second embodiment, the ability of a small molecule to displace a  $\mu$ -conopeptide from a sodium channel subtype is determined. In a third embodiment, the binding affinity of a small molecule for a sodium channel subtype is compared to the binding affinity of a  $\mu$ -conopeptide for the same sodium channel subtype.

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#### DETAILED DESCRIPTION OF THE PREFERRED EMBODIMENT

[0028] The present invention is to  $\mu$ -conopeptides, derivatives or pharmaceutically acceptable salts thereof. The present invention is further directed to the use of this peptide, derivatives thereof and pharmaceutically acceptable salts thereof for the treatment of disorders associated with voltage-gated sodium channels. Thus, the  $\mu$ -conopeptides or derivatives are useful as neuromuscular blocking agents, local anesthetic agents, analgesic agents and neuroprotective agents. The  $\mu$ -conopeptides are also useful for treating neuromuscular

disorders. The invention is further directed to nucleic acid sequences encoding the  $\mu$ -conopeptides and encoding propeptides, as well as the propeptides.

[0029] The present invention, in another aspect, relates to a pharmaceutical composition comprising an effective amount of an  $\mu$ -conopeptides, a murein thereof, an analog thereof, an active fragment thereof or pharmaceutically acceptable salts or solvates. Such a pharmaceutical composition has the capability of acting at voltage gated ion channels, and are thus useful for treating a disorder or disease of a living animal body, including a human, which disorder or disease is responsive to the partial or complete blockade of voltage gated ion channels of the central nervous system comprising the step of administering to such a living animal body, including a human, in need thereof a therapeutically effective amount of a pharmaceutical composition of the present invention.

[0030] The present invention is directed to the use of  $\mu$ -conopeptides as neuromuscular blockers and for treating neuromuscular disorders. As such, the  $\mu$ -conopeptides are useful for providing relaxation of muscle, for treating benign essential blepharospasm and other forms of focal dystonia and for anti-wrinkle use. Thus, in one aspect, the  $\mu$ -conopeptides are useful as neuromuscular blocking agents in conjunction with surgery or for intubation of the trachea by conventional parenteral administration e.g., intramuscular or intravenous administration in solution. In a second aspect, the  $\mu$ -conopeptides are useful as agents for treating neuromuscular disorders such as myofacial pain syndrome, chronic muscle spasm, dystonias and spasticity.

[0031] The primary factor detrimental to neurons in neurological disorders associated with deficient oxygen supply or mitochondrial dysfunction is insufficient ATP production relative to their requirement. As a large part of the energy consumed by brain cells is used for maintenance of the  $\text{Na}^+$  gradient across the cellular membrane, reduction of energy demand by down-modulation of voltage-gated  $\text{Na}^+$ -channels is one strategy for neuroprotection. In addition, preservation of the inward  $\text{Na}^+$  gradient may be beneficial because it is an essential driving force for vital ion exchanges and transport mechanisms such as  $\text{Ca}^{2+}$  homeostasis and neurotransmitter uptake. Thus, the  $\mu$ -conopeptides of the present invention are useful as neuroprotectants.

[0032] Thus, the pharmaceutical compositions of the present invention are useful as neuroprotectants, especially cerebroprotectants, neuromuscular blockers, analgesics (both as a local anesthetic and for general analgesia use) or adjuvants to general anesthetics. A "neurological disorder or disease" is a disorder or disease of the nervous system including, but

not limited to, global and focal ischemic and hemorrhagic stroke, head trauma, spinal cord injury, hypoxia-induced nerve cell damage as in cardiac arrest or neonatal distress or epilepsy. In addition, a "neurological disorder or disease" is a disease state and condition in which a neuroprotectant, anticonvulsant, analgesic and/or as an adjunct in general anesthesia may be indicated, useful, recommended or prescribed.

[0033] More specifically, the present invention is directed to the use of these compounds for reducing neurotoxic injury associated with conditions of hypoxia, anoxia or ischemia which typically follows stroke, cerebrovascular accident, brain or spinal cord trauma, myocardial infarct, physical trauma, drowning, suffocation, perinatal asphyxia, or hypoglycemic events.

10 The present invention is further directed to the use of these compounds for treating pain, including acute and chronic pain, such migraine, nociceptive and neuropathic pain.

[0034] A "neuroprotectant" is a compound capable of preventing the neuronal death associated with a neurological disorder or disease. An "analgesic" is a compound capable of relieving pain by altering perception of nociceptive stimuli without producing anesthesia or loss of consciousness. A "muscle relaxant" is a compound that reduces muscular tension. An "adjunct in general anesthesia" is a compound useful in conjunction with anesthetic agents in producing the loss of ability to perceive pain associated with the loss of consciousness.

20 [0035] The invention relates as well to methods useful for treatment of neurological disorders and diseases, including, but not limited to, global and focal ischemic and hemorrhagic stroke, head trauma, spinal cord injury, hypoxia-induced nerve cell damage such as in cardiac arrest or neonatal distress, epilepsy or other convulsive disorders without undesirable side effects.

[0036] Thus, in one embodiment, the invention provides a method of reducing/alleviating/ decreasing the perception of pain by a subject or for inducing analgesia in a subject comprising administering to the subject an effective amount of the pharmaceutical composition comprising a therapeutically effective amount of a  $\mu$ -conopeptide described herein or a pharmaceutically acceptable salt or solvate thereof. The pain may be acute, persistent, inflammatory or neuropathic pain. The  $\mu$ -conopeptides are useful as an analgesia for chronic and neuropathic pain states, such as trigeminal neuralgia, diabetic neuropathy, post-herpetic neuralgia, neuroma pain, phantom limb pain. These peptides are also useful for treating burn pain and as ocular anesthetics.

[0037] In a second embodiment, the invention provides a method of reducing/alleviating/decreasing the perception of pain by a subject or for inducing analgesia, particularly local analgesia, in a subject comprising administering to the subject an effective amount of the pharmaceutical composition comprising a therapeutically effective amount of a  $\mu$ -conopeptide described herein or a pharmaceutically acceptable salt or solvate thereof. These peptides are also useful for treating burn pain and as ocular anesthetics.

[0038] In a third embodiment, the invention provides a method of treating stroke, head or spinal cord trauma or injury, anoxia, hypoxia-induced nerve cell damage, ischemia, migraine, psychosis, anxiety, schizophrenia, inflammation, movement disorder, epilepsy, any other convulsive disorder or in the prevention of the degenerative changes connected with the same in a subject comprising administering to the subject an effective amount of the pharmaceutical composition comprising a therapeutically effective amount of a  $\mu$ -conopeptide described herein or a pharmaceutically acceptable salt or solvate thereof.

[0039] In a fourth embodiment, the invention provides a method for providing a neuromuscular block or for treating neuromuscular disorders, such as methods for providing relaxation of muscle, for treating benign essential blepharospasm and other forms of focal dystonia and for anti-wrinkle use. Thus, in one aspect, the  $\mu$ -conopeptides are useful as neuromuscular blocking agents in conjunction with surgery or for intubation of the trachea by conventional parenteral administration e.g., intramuscular or intravenous administration in solution. In a second aspect, the  $\mu$ -conopeptides are useful as agents for treating neuromuscular disorders such as myofacial pain syndrome, chronic muscle spasm, dystonias and spasticity.

[0040] The present invention is also directed to the use of  $\mu$ -conopeptides for screening small molecule libraries to identify small molecules that are selective blocking agents at specific sodium channel subtypes expressed in mammalian systems. In one embodiment, the blocking activity of a small molecule at a particular sodium channel subtype is compared to the blocking activity of a  $\mu$ -conopeptide at the same sodium channel subtype. In a second embodiment, the ability of a small molecule to displace a  $\mu$ -conopeptide from a sodium channel subtype is determined. In a third embodiment, the binding affinity of a small molecule for a sodium channel subtype is compared to the binding affinity of a  $\mu$ -conopeptide for the same sodium channel subtype.

[0041] The  $\mu$ -conopeptides described herein are sufficiently small to be chemically synthesized. General chemical syntheses for preparing the foregoing  $\omega$ -conotoxin peptides are

described hereinafter. Various ones of the  $\mu$ -conopeptides can also be obtained by isolation and purification from specific *Conus* species using the technique described in U.S. Patent Nos. 4,447,356 (Olivera et al., 1984); 5,514,774; 5,719,264; and 5,591,821, as well as in PCT published application WO 98/03189, the disclosures of which are incorporated herein by reference.

[0042] Although the  $\mu$ -conopeptides of the present invention can be obtained by purification from cone snails, because the amounts of  $\mu$ -conopeptides obtainable from individual snails are very small, the desired substantially pure  $\mu$ -conopeptides are best practically obtained in commercially valuable amounts by chemical synthesis using solid-phase strategy. For example, the yield from a single cone snail may be about 10 micrograms or less of  $\mu$ -conopeptides peptide. By "substantially pure" is meant that the peptide is present in the substantial absence of other biological molecules of the same type; it is preferably present in an amount of at least about 85% purity and preferably at least about 95% purity. Chemical synthesis of biologically active  $\mu$ -conopeptides peptides depends of course upon correct determination of the amino acid sequence.

[0043] The  $\mu$ -conopeptides can also be produced by recombinant DNA techniques well known in the art. Such techniques are described by Sambrook et al. (1989). A gene of interest (i.e., a gene that encodes a suitable  $\mu$ -conopeptides) can be inserted into a cloning site of a suitable expression vector by using standard techniques. These techniques are well known to those skilled in the art. The expression vector containing the gene of interest may then be used to transfet the desired cell line. Standard transfection techniques such as calcium phosphate co-precipitation, DEAE-dextran transfection or electroporation may be utilized. A wide variety of host/expression vector combinations may be used to express a gene encoding a conotoxin peptide of interest. Such combinations are well known to a skilled artisan. The peptides produced in this manner are isolated, reduced if necessary, and oxidized to form the correct disulfide bonds.

[0044] One method of forming disulfide bonds in the  $\mu$ -conopeptides of the present invention is the air oxidation of the linear peptides for prolonged periods under cold room temperatures or at room temperature. This procedure results in the creation of a substantial amount of the bioactive, disulfide-linked peptides. The oxidized peptides are fractionated using reverse-phase high performance liquid chromatography (HPLC) or the like, to separate peptides having different linked configurations. Thereafter, either by comparing these fractions with the

elution of the native material or by using a simple assay, the particular fraction having the correct linkage for maximum biological potency is easily determined. However, because of the dilution resulting from the presence of other fractions of less biopotency, a somewhat higher dosage may be required.

5 [0045] The peptides are synthesized by a suitable method, such as by exclusively solid-phase techniques, by partial solid-phase techniques, by fragment condensation or by classical solution couplings.

10 [0046] In conventional solution phase peptide synthesis, the peptide chain can be prepared by a series of coupling reactions in which constituent amino acids are added to the growing peptide chain in the desired sequence. Use of various coupling reagents, e.g., dicyclohexylcarbodiimide or diisopropylcarbonyldimidazole; various active esters, e.g., esters of N-hydroxyphthalimide or N-hydroxy-succinimide, and the various cleavage reagents, to carry out reaction in solution, with subsequent isolation and purification of intermediates, is well known classical peptide methodology. Classical solution synthesis is described in detail in the 15 treatise, "Methoden der Organischen Chemie (Houben-Weyl): Synthese von Peptiden," (1974). Techniques of exclusively solid-phase synthesis are set forth in the textbook, "Solid-Phase Peptide Synthesis," (Stewart and Young, 1969), and are exemplified by the disclosure of U.S. Patent 4,105,603 (Vale et al., 1978). The fragment condensation method of synthesis is exemplified in U.S. Patent 3,972,859 (1976). Other available syntheses are exemplified by U.S. 20 Patents No. 3,842,067 (1974) and 3,862,925 (1975). The synthesis of peptides containing  $\gamma$ -carboxyglutamic acid residues is exemplified by Rivier et al. (1987), Nishiuchi et al. (1993) and Zhou et al. (1996).

25 [0047] Common to such chemical syntheses is the protection of the labile side chain groups of the various amino acid moieties with suitable protecting groups which will prevent a chemical reaction from occurring at that site until the group is ultimately removed. Usually also common is the protection of an  $\alpha$ -amino group on an amino acid or a fragment while that entity reacts at the carboxyl group, followed by the selective removal of the  $\alpha$ -amino protecting group to allow subsequent reaction to take place at that location. Accordingly, it is common that, as a 30 step in such a synthesis, an intermediate compound is produced which includes each of the amino acid residues located in its desired sequence in the peptide chain with appropriate side-chain protecting groups linked to various ones of the residues having labile side chains.

[0048] As far as the selection of a side chain amino protecting group is concerned, generally one is chosen which is not removed during deprotection of the  $\alpha$ -amino groups during the synthesis. However, for some amino acids, e.g., His, protection is not generally necessary. In selecting a particular side chain protecting group to be used in the synthesis of the peptides, 5 the following general rules are followed: (a) the protecting group preferably retains its protecting properties and is not split off under coupling conditions, (b) the protecting group should be stable under the reaction conditions selected for removing the  $\alpha$ -amino protecting group at each step of the synthesis, and (c) the side chain protecting group must be removable, upon the completion of the synthesis containing the desired amino acid sequence, under reaction 10 conditions that will not undesirably alter the peptide chain.

[0049] It should be possible to prepare many, or even all, of these peptides using recombinant DNA technology. However, when peptides are not so prepared, they are preferably prepared using the Merrifield solid-phase synthesis, although other equivalent chemical syntheses known in the art can also be used as previously mentioned. Solid-phase synthesis is 15 commenced from the C-terminus of the peptide by coupling a protected  $\alpha$ -amino acid to a suitable resin. Such a starting material can be prepared by attaching an  $\alpha$ -amino-protected amino acid by an ester linkage to a chloromethylated resin or a hydroxymethyl resin, or by an amide bond to a benzhydrylamine (BHA) resin or paramethylbenzhydrylamine (MBHA) resin. Preparation of the hydroxymethyl resin is described by Bodansky et al. (1966). 20 Chloromethylated resins are commercially available from Bio Rad Laboratories (Richmond, CA) and from Lab. Systems, Inc. The preparation of such a resin is described by Stewart and Young (1969). BHA and MBHA resin supports are commercially available, and are generally used when the desired polypeptide being synthesized has an unsubstituted amide at the C-terminus. Thus, solid resin supports may be any of those known in the art, such as one having 25 the formulae -O-CH<sub>2</sub>-resin support, -NH BHA resin support, or -NH-MBHA resin support. When the unsubstituted amide is desired, use of a BHA or MBHA resin is preferred, because cleavage directly gives the amide. In case the N-methyl amide is desired, it can be generated from an N-methyl BHA resin. Should other substituted amides be desired, the teaching of U.S. Patent No. 4,569,967 (Kornreich et al., 1986) can be used, or should still other groups than the 30 free acid be desired at the C-terminus, it may be preferable to synthesize the peptide using classical methods as set forth in the Houben-Weyl text (1974).

[0050] The C-terminal amino acid, protected by Boc or Fmoc and by a side-chain protecting group, if appropriate, can be first coupled to a chloromethylated resin according to the procedure set forth in K. Horiki et al. (1978), using KF in DMF at about 60°C for 24 hours with stirring, when a peptide having free acid at the C-terminus is to be synthesized. Following the 5 coupling of the BOC-protected amino acid to the resin support, the  $\alpha$ -amino protecting group is removed, as by using trifluoroacetic acid (TFA) in methylene chloride or TFA alone. The deprotection is carried out at a temperature between about 0°C and room temperature. Other standard cleaving reagents, such as HCl in dioxane, and conditions for removal of specific  $\alpha$ -amino protecting groups may be used as described in Schroder & Lubke (1965).

10 [0051] After removal of the  $\alpha$ -amino-protecting group, the remaining  $\alpha$ -amino- and side chain-protected amino acids are coupled step-wise in the desired order to obtain the intermediate compound defined hereinbefore, or as an alternative to adding each amino acid separately in the synthesis, some of them may be coupled to one another prior to addition to the solid phase reactor. Selection of an appropriate coupling reagent is within the skill of the art. Particularly 15 suitable as a coupling reagent is N,N'-dicyclohexylcarbodiimide (DCC, DIC, HBTU, HATU, TBTU in the presence of HoBt or HoAt).

20 [0052] The activating reagents used in the solid phase synthesis of the peptides are well known in the peptide art. Examples of suitable activating reagents are carbodiimides, such as N,N'-diisopropylcarbodiimide and N-ethyl-N'-(3-dimethylaminopropyl)carbodiimide. Other activating reagents and their use in peptide coupling are described by Schroder & Lubke (1965) and Kapoor (1970).

25 [0053] Each protected amino acid or amino acid sequence is introduced into the solid-phase reactor in about a twofold or more excess, and the coupling may be carried out in a medium of dimethylformamide (DMF):CH<sub>2</sub>Cl<sub>2</sub> (1:1) or in DMF or CH<sub>2</sub>Cl<sub>2</sub> alone. In cases where intermediate coupling occurs, the coupling procedure is repeated before removal of the  $\alpha$ -amino protecting group prior to the coupling of the next amino acid. The success of the coupling reaction at each stage of the synthesis, if performed manually, is preferably monitored by the ninhydrin reaction, as described by Kaiser et al. (1970). Coupling reactions can be performed automatically, as on a Beckman 990 automatic synthesizer, using a program such as that 30 reported in Rivier et al. (1978).

[0054] After the desired amino acid sequence has been completed, the intermediate peptide can be removed from the resin support by treatment with a reagent, such as liquid

hydrogen fluoride or TFA (if using Fmoc chemistry), which not only cleaves the peptide from the resin but also cleaves all remaining side chain protecting groups and also the amino protecting group at the N-terminus if it was not previously removed to obtain the peptide in the form of the free acid. If Met is present in the sequence, the Boc protecting group is preferably 5 first removed using trifluoroacetic acid (TFA)/ethanedithiol prior to cleaving the peptide from the resin with HF to eliminate potential S-alkylation. When using hydrogen fluoride or TFA for cleaving, one or more scavengers such as anisole, cresol, dimethyl sulfide and methylethyl sulfide are included in the reaction vessel.

[0055] Cyclization of the linear peptide is preferably affected, as opposed to cyclizing 10 the peptide while a part of the peptido-resin, to create bonds between Cys residues. To effect such a disulfide cyclizing linkage, fully protected peptide can be cleaved from a hydroxymethylated resin or a chloromethylated resin support by ammonolysis, as is well known in the art, to yield the fully protected amide intermediate, which is thereafter suitably cyclized and deprotected. Alternatively, deprotection, as well as cleavage of the peptide from the above 15 resins or a benzhydrylamine (BHA) resin or a methylbenzhydrylamine (MBHA), can take place at 0°C with hydrofluoric acid (HF) or TFA, followed by oxidation as described above.

[0056] The peptides are also synthesized using an automatic synthesizer. Amino acids are sequentially coupled to an MBHA Rink resin (typically 100 mg of resin) beginning at the C-terminus using an Advanced Chemtech 357 Automatic Peptide Synthesizer. Couplings are 20 carried out using 1,3-diisopropylcarbodiimide in N-methylpyrrolidinone (NMP) or by 2-(1H-benzotriazole-1-yl)-1,1,3,3-tetramethyluronium hexafluorophosphate (HBTU) and diethylisopropylethylamine (DIEA). The FMOC protecting group is removed by treatment with a 20% solution of piperidine in dimethylformamide(DMF). Resins are subsequently washed with DMF (twice), followed by methanol and NMP.

[0057] Muteins, analogs or active fragments, of the foregoing conotoxin peptides are 25 also contemplated here. See, e.g., Hammerland et al. (1992). Derivative muteins, analogs or active fragments of the conotoxin peptides may be synthesized according to known techniques, including conservative amino acid substitutions, such as outlined in U.S. Patent Nos. 5,545,723 (see particularly col. 2, line 50--col. 3, line 8); 5,534,615 (see particularly col. 19, line 45--col. 30 22, line 33); and 5,364,769 (see particularly col. 4, line 55--col. 7, line 26), each herein incorporated by reference.

[0058] The  $\mu$ -conopeptides of the present invention are also useful to reduce neurotoxic injury associated with conditions of hypoxia, anoxia or ischemia which typically follows stroke, cerebrovascular accident, brain or spinal chord trauma, myocardial infarct, physical trauma, drownings, suffocation, perinatal asphyxia, or hypoglycemic events. To reduce neurotoxic injury, an  $\omega$ -conopeptide should be administered in a therapeutically effective amount to the patient within 24 hours of the onset of the hypoxic, anoxic or ischemic condition in order for the  $\mu$ -conopeptide to effectively minimize the CNS damage which the patient will experience.

[0059] The  $\mu$ -conopeptides of the present invention are further useful in controlling pain, e.g., as analgesic agents, and the treatment of migraine, acute pain or persistent pain. They can be used prophylactically or to relieve the symptoms associated with a migraine episode, or to treat acute or persistent pain. For these uses, an  $\mu$ -conopeptide is administered in a therapeutically effective amount to overcome or to ease the pain.

[0060] The  $\mu$ -conopeptides of the present invention are also useful as neuromuscular blockers and for treating neuromuscular disorders. They can be used for providing relaxation of muscle, for treating benign essential blepharospasm and other forms of focal dystonia and for anti-wrinkle use. Thus, in one aspect, the  $\mu$ -conopeptides are used as neuromuscular blocking agents in conjunction with surgery or for intubation of the trachea by conventional parenteral administration e.g., intramuscular or intravenous administration in solution. In a second aspect, the  $\mu$ -conopeptides are used as agents for treating neuromuscular disorders such as myofacial pain syndrome, chronic muscle spasm, dystonias and spasticity. For these uses, a  $\mu$ -conopeptide is administered in a therapeutically effective amount to relax muscle or provide a neuromuscular block.

[0061] Pharmaceutical compositions containing a compound of the present invention as the active ingredient can be prepared according to conventional pharmaceutical compounding techniques. See, for example, *Remington's Pharmaceutical Sciences*, 18th Ed. (1990, Mack Publishing Co., Easton, PA). Typically, an antagonistic amount of active ingredient will be admixed with a pharmaceutically acceptable carrier. The carrier may take a wide variety of forms depending on the form of preparation desired for administration, e.g., intravenous, oral, parenteral or intrathecally. For examples of delivery methods see U.S. Patent No. 5,844,077, incorporated herein by reference.

[0062] "Pharmaceutical composition" means physically discrete coherent portions suitable for medical administration. "Pharmaceutical composition in dosage unit form" means

physically discrete coherent units suitable for medical administration, each containing a daily dose or a multiple (up to four times) or a sub-multiple (down to a fortieth) of a daily dose of the active compound in association with a carrier and/or enclosed within an envelope. Whether the composition contains a daily dose, or for example, a half, a third or a quarter of a daily dose, will 5 depend on whether the pharmaceutical composition is to be administered once or, for example, twice, three times or four times a day, respectively.

[0063] The term "salt", as used herein, denotes acidic and/or basic salts, formed with inorganic or organic acids and/or bases, preferably basic salts. While pharmaceutically acceptable salts are preferred, particularly when employing the compounds of the invention as 10 medicaments, other salts find utility, for example, in processing these compounds, or where non-medicament-type uses are contemplated. Salts of these compounds may be prepared by art-recognized techniques.

[0064] Examples of such pharmaceutically acceptable salts include, but are not limited to, inorganic and organic addition salts, such as hydrochloride, sulphates, nitrates or phosphates 15 and acetates, trifluoroacetates, propionates, succinates, benzoates, citrates, tartrates, fumarates, maleates, methane-sulfonates, isothionates, theophylline acetates, salicylates, respectively, or the like. Lower alkyl quaternary ammonium salts and the like are suitable, as well.

[0065] As used herein, the term "pharmaceutically acceptable" carrier means a non-toxic, inert solid, semi-solid liquid filler, diluent, encapsulating material, formulation auxiliary of any 20 type, or simply a sterile aqueous medium, such as saline. Some examples of the materials that can serve as pharmaceutically acceptable carriers are sugars, such as lactose, glucose and sucrose, starches such as corn starch and potato starch, cellulose and its derivatives such as sodium carboxymethyl cellulose, ethyl cellulose and cellulose acetate; powdered tragacanth; malt, gelatin, talc; excipients such as cocoa butter and suppository waxes; oils such as peanut oil, 25 cottonseed oil, safflower oil, sesame oil, olive oil, corn oil and soybean oil; glycols, such as propylene glycol, polyols such as glycerin, sorbitol, mannitol and polyethylene glycol; esters such as ethyl oleate and ethyl laurate, agar; buffering agents such as magnesium hydroxide and aluminum hydroxide; alginic acid; pyrogen-free water; isotonic saline, Ringer's solution; ethyl alcohol and phosphate buffer solutions, as well as other non-toxic compatible substances used in 30 pharmaceutical formulations.

[0066] Wetting agents, emulsifiers and lubricants such as sodium lauryl sulfate and magnesium stearate, as well as coloring agents, releasing agents, coating agents, sweetening,

flavoring and perfuming agents, preservatives and antioxidants can also be present in the composition, according to the judgment of the formulator. Examples of pharmaceutically acceptable antioxidants include, but are not limited to, water soluble antioxidants such as ascorbic acid, cysteine hydrochloride, sodium bisulfite, sodium metabisulfite, sodium sulfite, 5 and the like; oil soluble antioxidants, such as ascorbyl palmitate, butylated hydroxyanisole (BHA), butylated hydroxytoluene (BHT), lecithin, propyl gallate, aloha-tocopherol and the like; and the metal chelating agents such as citric acid, ethylenediamine tetraacetic acid (EDTA), sorbitol, tartaric acid, phosphoric acid and the like.

[0067] For oral administration, the compounds can be formulated into solid or liquid 10 preparations such as capsules, pills, tablets, lozenges, melts, powders, suspensions or emulsions.

In preparing the compositions in oral dosage form, any of the usual pharmaceutical media may be employed, such as, for example, water, glycols, oils, alcohols, flavoring agents, preservatives, 15 coloring agents, suspending agents, and the like in the case of oral liquid preparations (such as, for example, suspensions, elixirs and solutions); or carriers such as starches, sugars, diluents, granulating agents, lubricants, binders, disintegrating agents and the like in the case of oral solid preparations (such as, for example, powders, capsules and tablets). Because of their ease in administration, tablets and capsules represent the most advantageous oral dosage unit form, in which case solid pharmaceutical carriers are obviously employed. If desired, tablets may be sugar-coated or enteric-coated by standard techniques. The active agent can be encapsulated to 20 make it stable to passage through the gastrointestinal tract while at the same time allowing for passage across the blood brain barrier. See for example, WO 96/11698.

[0068] For parenteral administration, the compound may be dissolved in a pharmaceutical carrier and administered as either a solution or a suspension. Illustrative of suitable carriers are water, saline, dextrose solutions, fructose solutions, ethanol, or oils of animal, vegetative or synthetic origin. The carrier may also contain other ingredients, for 25 example, preservatives, suspending agents, solubilizing agents, buffers and the like. When the compounds are being administered intrathecally, they may also be dissolved in cerebrospinal fluid.

[0069] A variety of administration routes are available. The particular mode selected will 30 depend of course, upon the particular drug selected, the severity of the disease state being treated and the dosage required for therapeutic efficacy. The methods of this invention, generally speaking, may be practiced using any mode of administration that is medically acceptable,

meaning any mode that produces effective levels of the active compounds without causing clinically unacceptable adverse effects. Such modes of administration include oral, rectal, sublingual, topical, nasal, transdermal or parenteral routes. The term "parenteral" includes subcutaneous, intravenous, epidural, irrigation, intramuscular, release pumps, or infusion.

5 [0070] For example, administration of the active agent according to this invention may be achieved using any suitable delivery means, including:

(a) pump (see, e.g., Luer & Hatton (1993), Zimm et al. (1984) and Ettinger et al. (1978));

(b), microencapsulation (see, e.g., U.S. Patent Nos. 4,352,883; 4,353,888; and 5,084,350);

10 (c) continuous release polymer implants (see, e.g., U.S. Patent No. 4,883,666);

(d) macroencapsulation (see, e.g., U.S. Patent Nos. 5,284,761, 5,158,881, 4,976,859 and 4,968,733 and published PCT patent applications WO92/19195, WO 95/05452);

(e) naked or unencapsulated cell grafts to the CNS (see, e.g., U.S. Patent Nos. 5,082,670 and 5,618,531);

15 (f) injection, either subcutaneously, intravenously, intra-arterially, intramuscularly, or to other suitable site; or

(g) oral administration, in capsule, liquid, tablet, pill, or prolonged release formulation.

20 [0071] In one embodiment of this invention, an active agent is delivered directly into the CNS, preferably to the brain ventricles, brain parenchyma, the intrathecal space or other suitable CNS location, most preferably intrathecally.

25 [0072] Alternatively, targeting therapies may be used to deliver the active agent more specifically to certain types of cell, by the use of targeting systems such as antibodies or cell specific ligands. Targeting may be desirable for a variety of reasons, e.g. if the agent is unacceptably toxic, or if it would otherwise require too high a dosage, or if it would not otherwise be able to enter the target cells.

30 [0073] The active agents, which are peptides, can also be administered in a cell based delivery system in which a DNA sequence encoding an active agent is introduced into cells designed for implantation in the body of the patient, especially in the spinal cord region. Suitable delivery systems are described in U.S. Patent No. 5,550,050 and published PCT Application Nos. WO 92/19195, WO 94/25503, WO 95/01203, WO 95/05452, WO 96/02286, WO 96/02646, WO 96/40871, WO 96/40959 and WO 97/12635. Suitable DNA sequences can

be prepared synthetically for each active agent on the basis of the developed sequences and the known genetic code.

[0074] Exemplary methods for administering such muscle relaxant compounds (e.g., so as to achieve sterile or aseptic conditions) will be apparent to the skilled artisan. Certain 5 methods suitable for administering compounds useful according to the present invention are set forth in Goodman and Gilman's *The Pharmacological Basis of Therapeutics*, 7th Ed. (1985). The administration to the patient can be intermittent; or at a gradual, continuous, constant or controlled rate. Administration can be to a warm-blooded animal (e.g. a mammal, such as a mouse, rat, cat, rabbit, dog, pig, cow or monkey); but advantageously is administered to a human 10 being. Administration occurs after general anesthesia is administered. The frequency of administration normally is determined by an anesthesiologist, and typically varies from patient to patient.

[0075] The active agent is preferably administered in an therapeutically effective amount. By a "therapeutically effective amount" or simply "effective amount" of an active 15 compound is meant a sufficient amount of the compound to treat the desired condition at a reasonable benefit/risk ratio applicable to any medical treatment. The actual amount administered, and the rate and time-course of administration, will depend on the nature and severity of the condition being treated. Prescription of treatment, e.g. decisions on dosage, timing, etc., is within the responsibility of general practitioners or specialists, and typically takes 20 account of the disorder to be treated, the condition of the individual patient, the site of delivery, the method of administration and other factors known to practitioners. Examples of techniques and protocols can be found in *Remington's Pharmaceutical Sciences*.

[0076] Dosage may be adjusted appropriately to achieve desired drug levels, locally or systemically. Typically the active agents of the present invention exhibit their effect at a dosage 25 range from about 0.001 mg/kg to about 250 mg/kg, preferably from about 0.01 mg/kg to about 100 mg/kg of the active ingredient, more preferably from about 0.05 mg/kg to about 75 mg/kg. A suitable dose can be administered in multiple sub-doses per day. Typically, a dose or sub-dose may contain from about 0.1 mg to about 500 mg of the active ingredient per unit dosage 30 form. A more preferred dosage will contain from about 0.5 mg to about 100 mg of active ingredient per unit dosage form. Dosages are generally initiated at lower levels and increased until desired effects are achieved. In the event that the response in a subject is insufficient at such doses, even higher doses (or effective higher doses by a different, more localized delivery

route) may be employed to the extent that patient tolerance permits. Continuous dosing over, for example 24 hours or multiple doses per day are contemplated to achieve appropriate systemic levels of compounds.

[0077] For the treatment of pain, if the route of administration is directly to the CNS, the 5 dosage contemplated is from about 1 ng to about 100 mg per day, preferably from about 100 ng to about 10 mg per day, more preferably from about 1  $\mu$ g to about 100  $\mu$ g per day. If administered peripherally, the dosage contemplated is somewhat higher, from about 100 ng to about 1000 mg per day, preferably from about 10  $\mu$ g to about 100 mg per day, more preferably from about 100  $\mu$ g to about 10 mg per day. If the conopeptide is delivered by continuous 10 infusion (e.g., by pump delivery, biodegradable polymer delivery or cell-based delivery), then a lower dosage is contemplated than for bolus delivery.

[0078] Advantageously, the compositions are formulated as dosage units, each unit being adapted to supply a fixed dose of active ingredients. Tablets, coated tablets, capsules, ampoules and suppositories are examples of dosage forms according to the invention.

15 [0079] It is only necessary that the active ingredient constitute an effective amount, i.e., such that a suitable effective dosage will be consistent with the dosage form employed in single or multiple unit doses. The exact individual dosages, as well as daily dosages, are determined according to standard medical principles under the direction of a physician or veterinarian for use humans or animals.

20 [0080] The pharmaceutical compositions will generally contain from about 0.0001 to 99 wt. %, preferably about 0.001 to 50 wt. %, more preferably about 0.01 to 10 wt.% of the active ingredient by weight of the total composition. In addition to the active agent, the pharmaceutical compositions and medicaments can also contain other pharmaceutically active compounds. Examples of other pharmaceutically active compounds include, but are not limited to, analgesic 25 agents, cytokines and therapeutic agents in all of the major areas of clinical medicine. When used with other pharmaceutically active compounds, the conopeptides of the present invention may be delivered in the form of drug cocktails. A cocktail is a mixture of any one of the compounds useful with this invention with another drug or agent. In this embodiment, a common administration vehicle (e.g., pill, tablet, implant, pump, injectable solution, etc.) would 30 contain both the instant composition in combination supplementary potentiating agent. The individual drugs of the cocktail are each administered in therapeutically effective amounts. A therapeutically effective amount will be determined by the parameters described above; but, in

any event, is that amount which establishes a level of the drugs in the area of body where the drugs are required for a period of time which is effective in attaining the desired effects.

[0081] The practice of the present invention employs, unless otherwise indicated, conventional techniques of chemistry, molecular biology, microbiology, recombinant DNA, genetics, immunology, cell biology, cell culture and transgenic biology, which are within the skill of the art. See, e.g., Maniatis *et al.*, 1982; Sambrook *et al.*, 1989; Ausubel *et al.*, 1992; Glover, 1985; Anand, 1992; Guthrie and Fink, 1991; Harlow and Lane, 1988; Jakoby and Pastan, 1979; *Nucleic Acid Hybridization* (B. D. Hames & S. J. Higgins eds. 1984); *Transcription And Translation* (B. D. Hames & S. J. Higgins eds. 1984); *Culture Of Animal Cells* (R. I. Freshney, Alan R. Liss, Inc., 1987); *Immobilized Cells And Enzymes* (IRL Press, 1986); B. Perbal, *A Practical Guide To Molecular Cloning* (1984); the treatise, *Methods In Enzymology* (Academic Press, Inc., N.Y.); *Gene Transfer Vectors For Mammalian Cells* (J. H. Miller and M. P. Calos eds., 1987, Cold Spring Harbor Laboratory); *Methods In Enzymology*, Vols. 154 and 155 (Wu *et al.* eds.), *Immunochemical Methods In Cell And Molecular Biology* (Mayer and Walker, eds., Academic Press, London, 1987); *Handbook Of Experimental Immunology*, Volumes I-IV (D. M. Weir and C. C. Blackwell, eds., 1986); Riott, *Essential Immunology*, 6th Edition, Blackwell Scientific Publications, Oxford, 1988; Hogan *et al.*, *Manipulating the Mouse Embryo*, (Cold Spring Harbor Laboratory Press, Cold Spring Harbor, N.Y., 1986).

### EXAMPLES

[0082] The present invention is described by reference to the following Examples, which are offered by way of illustration and are not intended to limit the invention in any manner. Standard techniques well known in the art or the techniques specifically described below were utilized.

#### EXAMPLE 1

##### Isolation of $\mu$ -Conopeptides

[0083] Crude venom was extracted from venom ducts (Cruz *et al.*, 1976), and the components were purified as previously described (Cartier *et al.*, 1996). The crude extract from venom ducts was purified by reverse phase liquid chromatography (RPLC) using a Vydac C<sub>18</sub>

semi-preparative column (10 x 250 mm). Further purification of bioactive peaks was done on a Vydac C<sub>18</sub> analytical column (4.6 x 220 mm). The effluents were monitored at 220 nm. Peaks were collected, and aliquots were assayed for activity. Throughout purification, HPLC fractions were assayed by means of intracerebral ventricular (i.c.v.) injection into mice (Clark et al., 5 1981).

[0084] The amino acid sequence of the purified peptides were determined by standard methods. The purified peptides were reduced and alkylated prior to sequencing by automated Edman degradation on an Applied Biosystems 477A Protein Sequencer with a 120A Analyzer (DNA/Peptide Facility, University of Utah) (Martinez et al., 1995; Shon et al., 1994).

10 [0085] In accordance with this method, the  $\mu$ -conopeptides described as "isolated" in Table 1 were obtained. These  $\mu$ -conopeptides, as well as the other  $\mu$ -conopeptides and the  $\mu$ -conopeptide precursors set forth in Table 1 are synthesized as described in U.S. Patent No. 5,670,622.

15

## EXAMPLE 2

### Isolation of DNA Encoding $\mu$ -Conopeptides

20 [0086] DNA coding for  $\mu$ -conopeptides was isolated and cloned in accordance with conventional techniques using general procedures well known in the art, such as described in Olivera et al. (1996). Alternatively, cDNA libraries was prepared from *Conus* venom duct using conventional techniques. DNA from single clones was amplified by conventional techniques using primers which correspond approximately to the M13 universal priming site and the M13 reverse universal priming site. Clones having a size of approximately 300-500 nucleotides were sequenced and screened for similarity in sequence to known  $\mu$ -conotoxins. The DNA sequences and encoded propeptide sequences are set forth in Table 1. DNA sequences coding for the 25 mature toxin can also be prepared on the basis of the DNA sequences set forth in Table 1. An alignment of the  $\mu$ -conopeptides of the present invention is set forth in Table 2.

### TABLE 1

#### DNA and Amino Acid Sequences of $\mu$ -Conopeptides and Precursors

30 Name: Ar3.1  
Species: arenatus  
Cloned: Yes

**DNA Sequence:**

CAAGAAGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTCTTGACCATCTG  
TATGCTTCTGTTCCCTTACTGCTCTCCGCTGGATGGGATCAACCTGCAGACCG  
5 ACCTGCAGAGCGTATGCAGGACACTTAACTGAGCATCATCCCCGTGATCC  
TGTCAAACGGTGTGCGAGAGGCCATGCAACATAGGATGCGTACCTGTTGTTAATG  
ACCAGCTTGTATCGCGGCCTCATCAAGCGAATAAGTAAACGATTGCAGT (SEQ  
ID NO:1)

**Translation:**

10 MMSKLGVLТИCMLLFPLTALPLGDQPADRPАЕRMQDDFITEHHPLFDPVKRCCERPC  
NIGCVPCC (SEQ ID NO:2)

**Toxin Sequence:**

Cys-Cys-Xaa1-Arg-Xaa3-Cys-Asn-Ile-Gly-Cys-Val-Xaa3-Cys-Cys-^ (SEQ ID NO:3)

15

Name: Ak3.1  
Species: atlanticus  
Cloned: Yes

**DNA Sequence:**

20 GGATCCATGATGTCTAAACTGGGAGTCTTGTGACCATCTGCTCTGTTCCAC  
TTACTGCTCTCCGCTGGATGAAGATCAACCGGTACACCGACCTGCAGAGCGTATGC  
AGGACATTCATCTGATCAACATCTCTTGTGATCTCATCAAACGGTGTGCGAGT  
25 TGCCATGCGGGCCAGGCTTTGCGTCCCTGTTGCTGACATCAATAACGTGTTGATG  
ACCAACTTCTCGAG (SEQ ID NO:4)

**Translation:**

30 GSMMMSKLGVLТИCMLLFPLTALPLDEDQPVHRPAЕRMQDISSDQHLFFDLIKRCCELPC  
GPGFCVPCC (SEQ ID NO:5)

**Toxin Sequence:**

Cys-Cys-Xaa1-Leu-Xaa3-Cys-Gly-Xaa3-Gly-Phe-Cys-Val-Xaa3-Cys-Cys-^ (SEQ ID NO:6)

35

Name: A3.1  
Species: aurisiacus  
Cloned: Yes

**DNA Sequence:**

40 CAAGAGGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTGACCATCTG  
TTGCTTCTGTTCCCTTACTGCTCTCCGATGGATGGAGATCAATCTGTAGACCGA  
CCTGAAGAGCGTATGCAGGACGACATTCTCATCTGAGCAGCATCCCTGTTAACAG  
AAAAGAATGTGTTGCGCGAAGGCCGAAATGCCAGCTATTCAGAAACAGTCA  
45 GATTGTCATTGTTAAATGACAACGTGTCGATGACCAACTCGTTATCACGACT  
AATGAATAAGTAAACGATTGCAGT (SEQ ID NO:7)

**Translation:**

MMSKLGVLLTICLLLFPPLTALPMGDQSVDRPEERMQDDISSEQHPLFNQKRMCCGEG  
RKCPsyFRNSQICHCC (SEQ ID NO:8)

**5 Toxin Sequence:**

Met-Cys-Cys-Gly-Xaa1-Gly-Arg-Lys-Cys-Xaa3-Ser-Xaa5-Phe-Arg-Asn-Ser-Gln-Ile-Cys-His-Cys-Cys-# (SEQ ID NO:9)

10 Name: A3.2  
Species: aurisiacus  
Cloned: Yes

**DNA Sequence:**

15 GGATCCATGATGTCTAAACTGGGAGTCTTGTGACCCTGTGTTGCTTCTGTTCCCC  
TTACTGCTTCCGATCGATGGAGATCAATCTGTAGACCGACCTGCAGAGCGTATGC  
AGGATGACATTTCATCTGAGCAGCAGTCGCTTGTCAATCAGAAAAGAAGGTGCTGC  
CGGTGGCCATGCCCGACAAATCGACGGTGAATTGTGGCTGTCCTGGATGA  
TAACCGTGTGATGACCAACTTCTCGAG (SEQ ID NO:10)

**20 Translation:**

GSMSKLGVLLTICLLLFPPLTALPDGDQSVDRPAERMQDDISSEQHRLFNQKRRCCRW  
PCPRQIDGEYCGCCLG (SEQ ID NO:11)

**25 Toxin Sequence:**

Cys-Cys-Arg-Xaa4-Xaa3-Cys-Xaa3-Arg-Gln-Ile-Asp-Gly-Xaa1-Xaa5-Cys-Gly-Cys-Cys-Leu-# (SEQ ID NO:12)

30 Name: A3.3  
Species: aurisiacus  
Cloned: Yes

**DNA Sequence:**

35 GGATCCATGATGTCTAAACTGGGAGTCTTGTGACCCTGTCTACTTCTGTTCCCC  
TTACTGCTTCCGATGGATGGAGATCAACCTGCAGACCAACCTGCAGATCGTATGC  
AGGACGACATTTCATCTGAGCAGTATCCCTTGTGATAAGAGACAAAAGTGTGCA  
CTGGGAAGAAGGGGTATGCTCCGGCAAAGCATGCAAAATCTCAAATGTTGCTCT  
GGACGATAACGTGTGATGACCAACTTCTCGAG (SEQ ID NO:13)

**40 Translation:**

GSMSKLGVLLTICLLLFPPLTAFPMGDQPADQPADRMQDDISSEQYPLFDKRQKCT  
GKKGSCSGKACKNLKCCSGR (SEQ ID NO:14)

**45 Toxin Sequence:**

Xaa2-Lys-Cys-Cys-Thr-Gly-Lys-Lys-Gly-Ser-Cys-Ser-Gly-Lys-Ala-Cys-Lys-Asn-Leu-Lys-Cys-Cys-Ser-# (SEQ ID NO:15)

5      **Name:**      A3.4  
      **Species:**      aurisiacus  
      **Cloned:**      Yes

**DNA Sequence:**  
GGATCCATGATGTCTAAACTGGGAGTCTTGTGACCATCTGTCTGCTTCTGTTCCAC  
TTACTGCTGTTCCGCTGGATGGAGATCAACCTCTAGACCGACACGCGAGCGTATGC  
10     ATGATGGCATTTCACCTAAACGCCATCCCTGGTTGATCCCGTCAAACGGTGTGCA  
AGGTGCAATGCGAGTCTGACCCCTGTTGCTAACGTGTTGATGACCAACTTCTC  
GAG (SEQ ID NO:16)

15     **Translation:**  
GSMMMSKLGVLLTICLLLFPPLAVPLDGDQPLDRHAERMHDGISPKRHPWFDPVKRCK  
VQCESCTPCC (SEQ ID NO:17)

20     **Toxin Sequence:**  
Cys-Cys-Lys-Val-Gln-Cys-Xaa1-Ser-Cys-Thr-Xaa3-Cys-Cys-^ (SEQ ID NO:18)

25     **Name:**      Bn3.1  
      **Species:**      bandanus  
      **Cloned:**      Yes

30     **DNA Sequence:**  
GGATCCATGATGTCTAAACTGGGAGTCTTGTGACCATCTGTATGCTTCTGTTCCCC  
TCACTGCTTCCGATGGATGGAGATCAACCTGCAGACCGACCTGCAGAGCGTAGT  
CAGGACGTTTCATCTGAACAGCATCCCTGTTGATCCCGTCAAACGGTGTGCAAC  
TGGCCATGCTCCATGGGATGCATCCCTGTTGCTACTATTAAATAACGTGTTGATGAC  
CAACTTCTCGAG (SEQ ID NO:19)

35     **Translation:**  
GSMMMSKLGVLLTICMLLFPLTALPMGDQPADRPAERSQDVSSSEQHPLFDPVKRCCNW  
PCSMGCIIPCCYY (SEQ ID NO:20)

40     **Toxin Sequence:**  
Cys-Cys-Asn-Xaa4-Xaa3-Cys-Ser-Met-Gly-Cys-Ile-Xaa3-Cys-Cys-Xaa5-Xaa5-^ (SEQ ID  
NO:21)

45     **Name:**      Bt3.1  
      **Species:**      betulinus  
      **Cloned:**      Yes

**DNA Sequence:**  
CAAGAGGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTGACCTCTG

5 TCTGCTTCTGTTCCCTGACTGCTCTCCGCTGGATGAAGATCAACCTGCAGACCG  
 ACCTGCAGAGCGTATGCAGGACATTCATCTGAACAGCATCCCTGTTGATCCCGT  
 CAAACGGTGTGCGAATTGCCATGCCATGGATGCGTCCCTGTTGCTGGCCTTAATA  
 ACGTGTGGATGACCAACTGTGTATCACGGCCACGTCAAGTGTCTAATGAATAAGT  
 AAAATGATTGCAGT (SEQ ID NO:22)

**Translation:**

MMSKLGVLTFCLLFPLTALPLDEDQPADRPAERMQDISSEQHPLFDPVKRCCELPCH  
 GCVPCCWP (SEQ ID NO:23)

10

**Toxin Sequence:**

Cys-Cys-Xaa1-Leu-Xaa3-Cys-His-Gly-Cys-Val-Xaa3-Cys-Cys-Xaa4-Xaa3-^ (SEQ ID NO:24)

15

Name: Bt3.2  
 Species: betulinus  
 Cloned: Yes

**DNA Sequence:**

20 CAAGAGGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTGACCTCTG  
 TCTGCTTCTGTTCCCTGACTGCTCTCCGCTGGATGAAGATCAACCTGCAGACCG  
 ACATGCAGAGCGTATGCAGGACATTCACCTGAACAGCATCCCTGTTGATCCCGT  
 CAAACGGTGTGCGGGCTGCCATGCAATGGATGCGTCCCTGTTGCTGGCCTTCATA  
 ACGTGTGGACGACCAACTTGTATCACGGCCACGTCAAGTGTCTGATGAATAAGTA  
 25 AAACGATTGCAGT (SEQ ID NO:25)

**Translation:**

MMSKLGVLTFCLLFPLTALPLDEDQPADRHAERMQDISPEQHPSFDPVKRCGLPCN  
 GCVPCCWPS (SEQ ID NO:26)

30

**Toxin Sequence:**

Cys-Cys-Gly-Leu-Xaa3-Cys-Asn-Gly-Cys-Val-Xaa3-Cys-Cys-Xaa4-Xaa3-Ser-^ (SEQ ID NO:27)

35

Name: Bt3.3  
 Species: betulinus  
 Cloned: Yes

40

**DNA Sequence:**

CAAGAGGGATCGATAGCAGTTCATGATGTTAAACTGGGAGTCTTGTGACCATCTA  
 TATGCTTCTGTTCCCTTACTGCTCTCCGCTGGATGGAGATCAACCTGCAGACCAA  
 CCTCTAGAGCGCATGCAGTATGACATGTACGTGCGAGTGAATCCCTGGTTGATCCC  
 45 GTCAAAAGGTGCTGCTGAGGAACGTGCGAGTATGCATCCCTGTTGCCCGAATTGG  
 CCAGCTTGTGATTATCGCGGCCAAGAGTCTAATGAATAAGTAAAACGATTGCAGT (SEQ ID NO:28)

**Translation:**

MMFKLGVLITIYMLLFPTALPLDGDQPADQPLERMQYDMLRAVNPWFDPVKRCCSR  
NCAVCIPCCPNWPA (SEQ ID NO:29)

5

**Toxin Sequence:**

Cys-Cys-Ser-Arg-Asn-Cys-Ala-Val-Cys-Ile-Xaa3-Cys-Cys-Xaa3-Asn-Xaa4-Xaa3-Ala-<sup>5</sup> (SEQ ID NO:30)

10

Name: Bu3.1  
Species: bullatus  
Cloned: Yes

15

**DNA Sequence:**

CAAGAAGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTGACCATCTG  
TCTGCTTCTGTTCCCTTTGCTCTTCCGCAGGATGGAGATCAACCTGCAGACCGA  
CCTGCAGAGCGTATGCAGGACATTCATCTGAGCAGAATTCCCTGCTTGAGAA  
GAGAGTTACTGACAGGTGCTGCAAAGGGAAGAGGGAATGCGGCAGATGGTGCAGA  
20 GATCACTCGCGTTGTTGCGGTCGACGATAAGCTGTTGATGACCAGCTTGTTATCAC  
GGCTACATCAAGTGTCTAGTGAATAAGTAAAATGATTGCAGT (SEQ ID NO:31)

**Translation:**

MMSKLGVLITICLLFPLFALPQDGDQPADRPAERMQDDISSEQNSLLEKRVTDRCCKG  
25 KRECGRWCRDHSRCCGRR (SEQ ID NO:32)

**Toxin Sequence:**

Val-Thr-Asp-Arg-Cys-Cys-Lys-Gly-Lys-Arg-Xaa1-Cys-Gly-Arg-Xaa4-Cys-Arg-Asp-His-Ser-  
Arg-Cys-Cys-# (SEQ ID NO:33)

30

Name: Bu3.1A  
Species: bullatus  
Cloned: Yes

35

**DNA Sequence:**

CAAGAGGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTGACCATCTG  
TCTGCTTCTGTTCCCTTTGCTCTTCCGCAGGATGGAGATCAACCTGCAGACCGA  
CCTGCAGAGCGTATGCAGGATGACATTCATCTGAGCAGAATTCCCTGCTTGAGAA  
40 GAGAGTTGGTACAGGTGCTGCAAAGGGAAGAGGGGGTGCGGCAGATGGTGCAGA  
GATCACTCACGTTGTTGCGGTCGACGATAACGTGTTGATGACCAGCTTGTTATCAC  
GGCTACATCAAGTGTCTAGTGAATAAGTAAAACGATTGCAGT (SEQ ID NO:34)

**Translation:**

45

MMSKLGVLITICLLFPLFALRQDGDQPADRPAERMQDDISSEQNPILLEKRVGDRCKK  
GKRGCGRWCRDHSRCCGRR (SEQ ID NO:35)

**Toxin Sequence:**

Val-Gly-Asp-Arg-Cys-Cys-Lys-Gly-Lys-Arg-Gly-Cys-Gly-Arg-Xaa4-Cys-Arg-Asp-His-Ser-Arg-Cys-Cys-# (SEQ ID NO:36)

5

**Name:** Bu3.2  
**Species:** bullatus  
**Cloned:** Yes

10 **DNA Sequence:**

CAAGAAGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTGACCATCTG  
 TCTGCTTCTGTTCCCTTTGCTCTTCCGCAGGATGGAGATCAACCTGCAGACCGA  
 CCTGCAGAGCGTATGCAGGACGACATTCTATCTGAGCAGAATCCCTGCTTGAGAA  
 GAGAGTTGGTGAAGGTGCTGCAAAAACGGGAAGAGGGGGTGCAGATGGTGC  
 15 AGAGATCACTCACGTTGCGGTCGACGATAACGTGTTGATGACCGAGGCTTCGT  
 TATCACGGCTACATCAAGTGTCTAGTGAATAAGTAAAACGATTGCAGT (SEQ ID  
 NO:37)

**Translation:**

20 MMSKLGVLLTICLLLFPFLALPQDGDQPADRPAERMQDDISSEQNPLLEKRVGERCCKN  
 GKRGCGRWCRDHSRCCGRR (SEQ ID NO:38)

**Toxin Sequence:**

25 Val-Gly-Xaa1-Arg-Cys-Cys-Lys-Asn-Gly-Lys-Arg-Gly-Cys-Gly-Arg-Xaa4-Cys-Arg-Asp-His-Ser-Arg-Cys-Cys-# (SEQ ID NO:39)

30 **Name:** Bu3.3  
**Species:** bullatus  
**Cloned:** Yes

**DNA Sequence:**

CAAGAGGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTGACCATCTG  
 TCTGCTTCTGTTCCCTTTGCTCTTCCGCAGGACGGAGATCAACCTGCAGACCG  
 35 ACCTGCAGAGCGTATGCAGGACGACCTTCATCTGAGCAGCATCCCTGTTGAGAA  
 GAGAATTGTTGACAGGGTCTGCAACAAAGGGAACGGGAAGAGGGGGTGCAGCAGA  
 TGGTGCAGAGATCACTCACGTTGCGGTCGACGATGAACGTGTTGATGACCGAGG  
 CTTTGGTTATCACGGCTACATCAAGTGTCTAGTGAATAAGTAAAACGATTGCAGT  
 (SEQ ID NO:40)

40

**Translation:**

MMSKLGVLLTICLLLFPFLALPQDGDQPADRPAERMQDDLSSEQHPLFEKRVDRCCNK  
 GNGKRGCSRWC RDHSRCCGRR (SEQ ID NO:41)

45 **Toxin Sequence:**

Ile-Val-Asp-Arg-Cys-Cys-Asn-Lys-Gly-Asn-Gly-Lys-Arg-Gly-Cys-Ser-Arg-Xaa4-Cys-Arg-Asp-His-Ser-Arg-Cys-Cys-# (SEQ ID NO:42)

5      Name:      Bu3.4  
Species:      bullatus  
Cloned:      Yes

DNA Sequence:  
CAAGAAGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTGACCATCTG  
10     TCTGCTTCTGTTCCCCTTTGCTCTTCCGAGGATGGAGATCAACCTGCAGACCGA  
CCTGCTGAGCGTATGCAGGACGACATTCTGAGCGGAATCCCTGTTGAGAAG  
AGCGTTGGTTATATTGCTGCCGACCCAAACCCAACGGGCAGATGATGTGCGACAG  
15     ATGGTGCAGAAAAAAACTCACGTTGCGGGTCGACGATAATGTGTTGATGACCAGC  
TTTGTATCAAGGCTACATCAAGTATCTAGTGAATAAGTAAAACGATTGCAGT (SEQ  
ID NO:43)

15     Translation:  
MMSKLGVLLTICLLLFPFLALPQDGDQPADRPAERMQDDISERNPLFEKSVGLYCCRP  
KPNGQMMCDRWCEKNSRCCGRR (SEQ ID NO:44)

20     Toxin Sequence:  
Val-Gly-Leu-Xaa5-Cys-Cys-Arg-Xaa3-Lys-Xaa3-Asn-Gly-Gln-Met-Met-Cys-Asp-Arg-Xaa4-  
Cys-Xaa1-Lys-Asn-Ser-Arg-Cys-Cys-# (SEQ ID NO:45)

---

25     Name:      Bu3.5  
Species:      bullatus  
Cloned:      Yes

30     DNA Sequence:  
CAAGAAGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTGACCATCTG  
TCTGCTTCTGTTCCCCTTACTGCTCTCCGATGGATGGAGATCAATCTGTAGACCGA  
35     CCTGCAGAACGTATGCAGGACGACCTTCTGAGCAGCATCCCTGTTGTCAG  
AAAAGAAGGTGTTGCGCGAAGGCTTGACATGCCAGATATTGGAAAAACAGTCA  
GATTGTCCTGTTGTTAAATGACAACGTGTCGATGACCAACTCGGTATCACGACT  
ACGCCAAGTGTCTAATGAATAAGTAAAACGATTGCAGT (SEQ ID NO:46)

35     Translation:  
MMSKLGVLLTICLLLFPALTALPMGDQSVDRPAERMQDDLSSEQHPLFVQKRRCCGEG  
40     LTCPRYWKNSQICACC (SEQ ID NO:47)

45     Toxin Sequence:  
Arg-Cys-Cys-Gly-Xaa1-Gly-Leu-Thr-Cys-Xaa3-Arg-Xaa5-Xaa4-Lys-Asn-Ser-Gln-Ile-Cys-Ala-  
Cys-Cys-^ (SEQ ID NO:48)

Name: Bu3.5A  
Species: bullatus  
Cloned: Yes

5 **DNA Sequence:**

CAAGAGGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTGACCATCTG  
TCTGCTTCTGTTCCCTTTGCTCTTCCGCAGGATGGAGATCAACCTGCAGACCGA  
CCTGCTGAGCGTATGCAGGACATTCATCTGAGCAGGATCCCTGTTGTCAG  
AAAAGAAGGTGTTGCGCGAAGGCTTGACATGCCAGATATTGGAAAAACAGTCA  
10 GATTGTGCTTGTGTTAAATGACAACGTGTGATGACCAACTCGGTATCACGACTA  
CGCCAAGTGTCTAATGAATAAGTAAAACGATTGCAGT (SEQ ID NO:49)

Translation:

MMSKLGVLVTICLLLFPLFALPQDGDPADRPAERMQDDISSEQDPLFVQKRRCCGEGL  
15 TCPRYWKNSQICACC (SEQ ID NO:50)

Toxin Sequence:

Arg-Cys-Cys-Gly-Xaa1-Gly-Leu-Thr-Cys-Xaa3-Arg-Xaa5-Xaa4-Lys-Asn-Ser-Gln-Ile-Cys-Ala-  
Cys-Cys-<sup>^</sup> (SEQ ID NO:51)

20

Name: Cp3.1  
Species: capitaneus  
Cloned: Yes

25 **DNA Sequence:**

GGATCCATGATGTCTAAACTGGGAGTCTTGGTGACCATCTGCCTGCTTCTGTTCCC  
CTTGCTGCTTCCACTGGATGGAAATCAACCTGCAGACCACCTGCAAAGCGTACG  
CAAGATGACAGTTCAGCTGCCCTGATCAATACTGGATTGATCATTCCCATTCTGC  
30 TGCAGGGACTCGGGTAAGATTGTGTTGGTGTGCCGTAACGTGTTGATGACCAA  
CTTCTCGAG (SEQ ID NO:52)

Translation:

GSMMMSKLGVLVTICLLLFPLAAFPLDGNQPADHPAKRTQDDSSAALINTWIDHSHSCCR  
35 DCGEDCVGCCR (SEQ ID NO:53)

Toxin Sequence:

Ser-Cys-Cys-Arg-Asp-Cys-Gly-Xaa1-Asp-Cys-Val-Gly-Cys-Cys-Arg-<sup>^</sup> (SEQ ID NO:54)

40

Name: Ca3.1  
Species: characteristicus  
Cloned: Yes

45 **DNA Sequence:**

CAAGAGGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTGACCATCTG  
TCTGCTTCTGTTCCCTTACTGCTCTCCAATGGATGGAGATCAACCTGCAGACCA

ACCTGCAGATCGTATGCAGGACGACATTGATCTGAGCAGTATCCCTGTTGATAT  
 GAGAAAAAGGTGTTGCCGCCCCGGCGGTATGCCCGTATTTGAGAGACAATT  
 TTATTGTGGTTGTTAAATGACAACGTGTCGATGACCAACTTCATTATCACGAC  
 TACGCCAAGTGTCTAATGAATAAGTAAAATGATTGCAGT (SEQ ID NO:55)

5

**Translation:**

MMSKLVLLTICLLFPLTALPMGDQPADQPADRMQDDISSEQYPLFDMRKRCGPG  
 GSCPVYFRDNFICGCC (SEQ ID NO:56)

10

**Toxin Sequence:**

Cys-Cys-Gly-Xaa3-Gly-Gly-Ser-Cys-Xaa3-Val-Xaa5-Phe-Arg-Asp-Asn-Phe-Ile-Cys-Gly-Cys-  
 Cys-^ (SEQ ID NO:57)

15

**Name:** Ca3.2

**Species:** characteristicus

**Cloned:** Yes

**DNA Sequence:**

20

CAAGAGGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTGTTGACCATCTG  
 TCTGCTTCTGTTCCCTTACTGCTCTTCCATGGATGGAGATGAACCTGCAAACCG  
 ACCTGTCGAGCGTATGCAGGACAACTTCATCTGAGCAGTATCCCTGTTGAGAA  
 GAGACGAGATTGTTGCACTCCGCCAAGAAATGCAAAGACCGACAATGCAAACCCC  
 AGAGATGTTGCGCTGGACGATAACGTGTTGATGACCAACTTATCACGGCTACGTCA  
 25 AGTGTAGTGAATAAGTAAAATGATTGCAGT (SEQ ID NO:58)

**Translation:**

MMSKLVLLTICLLFPLTALPMGDDEPANRPVERMQDNISSEQYPLFEKRRDCCTPPK  
 KCKDRQCKPQRCCAGR (SEQ ID NO:59)

30

**Toxin Sequence:**

Arg-Asp-Cys-Cys-Thr-Xaa3-Xaa3-Lys-Lys-Cys-Lys-Asp-Arg-Gln-Cys-Lys-Xaa3-Gln-Arg-  
 Cys-Cys-Ala-# (SEQ ID NO:60)

35

**Name:** Ca3.3

**Species:** characteristicus

**Cloned:** Yes

40

**DNA Sequence:**

CAAGAGGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTGTTGACCATCTG  
 TCTGCTTCTGTTCCCTTACTGCTCTTCCACTGGATGGAGATCAACCTGCAAGATCAA  
 TCTGCAGAGCGACCTGCAGAGCGTACGCAGGACGACATTGAGCAGCATCCGTTATA  
 TGATCCGAAAAGAAGGTGTTGCCGTATCCATGCCCGACAGCTGCCACGGATCTG  
 45 CTGCTATAAGTGATAACATGTTGATGGCCAGCTTGTATCACGGCCACGTCAAGTG  
 TCTTAATGAATAAGTAAAACGATTGCAGT (SEQ ID NO:61)

**Translation:**

MMSKLGVLLTICLLLPLTALPLDGDQPADQSAERPAERTQDDIQQHPLYDPKRRCCRY  
PCPDSCHGSCCYK (SEQ ID NO:62)

**5 Toxin Sequence:**

Arg-Cys-Cys-Arg-Xaa5-Xaa3-Cys-Xaa3-Asp-Ser-Cys-His-Gly-Ser-Cys-Cys-Xaa5-Lys-^ (SEQ ID NO:63)

10 Name: Ca3.4  
Species: characteristicus  
Cloned: Yes

**DNA Sequence:**

15 CAAGAGGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGCCTGTTGACCATCT  
GTCTACTTCTGTTTCCCTTAAGTGTGTTCCGCTGGATGGAGATCAACATGCAGACC  
AACCTGCACAGCGTCTGCAGGACCGCATTCAACTGAAGATCATCCCTTATTGATC  
CCAACAAACGGTGTGCCCCGGTGGCATGCAACATGGGATGCAAGCCTGTTGT  
GGATGACCAGCTTGTATCGCGGTCTCATGAAGTGTCTTAATGAATAAGTAAAAT  
20 GATTGCAGT (SEQ ID NO:64)

**Translation:**

MMSKLGALLTICLLLFSLTAVPLDGDQHADQPAQRLQDRIPTEHPLFDPNKRCCPPVA  
CNMGCKPCCG (SEQ ID NO:65)

25 **Toxin Sequence:**  
Cys-Cys-Xaa3-Xaa3-Val-Ala-Cys-Asn-Met-Gly-Cys-Lys-Xaa3-Cys-Cys-# (SEQ ID NO:66)

30 Name: Ca3.5  
Species: characteristicus  
Cloned: Yes

**DNA Sequence:**

35 CAAGAGGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGCCTGTTGACCATCT  
GTCTACTTCTGTTTCCCTTAAGTGTGTTCCGCTGGATGGAGATCAACATGCAGACC  
AACCTGCAGAGCGTCTGCATGACCGCCTTCAACTGAAAATCATCCCTTATATGATC  
CCGTCAAACGGTGTGCGATGATTGGAAATGCGACTATTCTGCTGGCCTGCTGTA  
TGTTTGGATAACCTTGTATCGCGGCCTCATCAAGTGTCTTAATGAATAAGTAAAAC  
40 GATTGCAGT (SEQ ID NO:67)

**Translation:**

MMSKLGALLTICLLLFSLTAVPLDGDQHADQPAERLHDRLPTEHPLYDPVKRCCDDSE  
CDYSCWPCCMFG (SEQ ID NO:68)

45 **Toxin Sequence:**  
Cys-Cys-Asp-Asp-Ser-Xaa1-Cys-Asp-Xaa5-Ser-Cys-Xaa4-Xaa3-Cys-Cys-Met-Phe-# (SEQ ID

NO:69)

5      **Name:**      Ca3.6  
**Species:**      characteristicus  
**Cloned:**      Yes

**DNA Sequence:**

GGATCCATGATGTCTAAACTGGGAGTCITGTTGACCATCTGCTGCTTCTGTTCCCC  
10     TTACTGCTGTTCCGCTGGATGGAGATCAACCTGCAGACCGACCTGCAGAGCGTAAG  
CAGGACGTTTCATCTGAACAGCATTCCCTTGTGATCCCGTCAAACGGTGTGCCGC  
CGGTGTTACATGGGATGCATCCCTTGTGCTTTAACGTGTTGATGACCAACTTCTC  
GAG (SEQ ID NO:70)

15      **Translation:**

GSMMMSKLGVLLTICLLLFPPLAVPLGDQPADRPAERKQDVSEQHPFFDPVKRCCRRC  
YMGCIPCCF (SEQ ID NO:71)

**Toxin Sequence:**

20      Cys-Cys-Arg-Arg-Cys-Xaa5-Met-Gly-Cys-Ile-Xaa3-Cys-Cys-Phe-^ (SEQ ID NO:72)

25      **Name:**      Cr3.1  
**Species:**      circumcisus  
**Cloned:**      Yes

**DNA Sequence:**

CAAGAAGGATCGATAGCAGTTCATGATGTCTAAACTGGGGTATTGTTGACCATCT  
GTCTGCTTCTGTTCCCTTACTGCTCTCCAATGGATGGAGATCAACCTGCAGACC  
30     AACCTGCAGATCGTATGCAGGACGACATTTCATCTGAGCAGATCCCTGTTGATA  
AGAGACGAAAGTGTGCGGCAAAGACGGGCCATGCCCAAATATTCAAAGACAAT  
TTTATTGTGGTTGTTAAATGACAACGTGTCGATGACCAACTCGTTATCACGAT  
TCGCCAAGTGTCTTAATGAATAAGTAAAATGATTGCAGT (SEQ ID NO:73)

35      **Translation:**

MMSKLGVLLTICLLLFPPLTALPMGDQPADQPADRMQDDISSEQYPLFDKRRKCCGKD  
GCPCKYFKDNFICGCC (SEQ ID NO:74)

**Toxin Sequence:**

40      Arg-Lys-Cys-Cys-Gly-Lys-Asp-Gly-Xaa3-Cys-Xaa3-Lys-Xaa5-Phe-Lys-Asp-Asn-Phe-Ile-Cys-  
Gly-Cys-Cys-^ (SEQ ID NO:75)

45      **Name:**      Da3.1  
**Species:**      dalli  
**Cloned:**      Yes

**DNA Sequence:**

CAAGAGGGATCGATAGCAGTTCATGATGTCATAAAGCTGGAGCCTGTTGACCATCT  
GTCTACTTCTGTTCCCTAACTGCTGTTCCGCTGGATGGAGATCAACATGCAGACC  
AACCTGCAGAGCGTCTGCAGGACCGCCTCCAAGTAAAATCATCCCTTATATGATC  
5 CCGTCAAACGGTGTGCGATGATCGGAATGCGACTATTCTGCTGGCCTGCTGTA  
TTTATCATAACCTTGTATCGCGGCTCATCAAGTGTCAAATGAATAAGTAAAAT  
GATTGCAGT (SEQ ID NO:76)

**Translation:**

10 MMSKLGALLTICLLLFSLTAVPLDGDQHADQPAERLQDRLPTENHPLYDPVKRCCDDSE  
CDYSCWPCCILS (SEQ ID NO:77)

**Toxin Sequence:**

15 Cys-Cys-Asp-Asp-Ser-Xaa1-Cys-Asp-Xaa5-Ser-Cys-Xaa4-Xaa3-Cys-Cys-Ile-Leu-Ser-^ (SEQ  
ID NO:78)

Name: Da3.2

Species: dalli

20 Cloned: Yes

**DNA Sequence:**

CAAGAGGGATCGATAGCAGTTCATGATGTCATAAAGCTGGAGCCTGTTGACCATTG  
TCTACTTCTGTTCCCTAACTGCTGTTCCACTGGATGGAGATCAGCCTGCAGACCG  
25 ACCTGCAGAGCGTATGCAGGACGGCATTCATCTGAACATCATCCATTGGATTC  
CGTAAAAAGAAAACAACAGTGTGCCCCGGTGGCATGCAACATGGGATGCGAGC  
CTTGTGTGGATGACCAGCTTGTATCGCGGCTCATGAAGTGTCTAATGAATAAG  
TAAAACGATTGCAGT (SEQ ID NO:79)

30 **Translation:**

MMSKLGVLLTICLLLPLTAVPLDGDQPADRPAERMQDGISSEHPFFDSVKKKQQCCP  
PVACNMGCEPCCG (SEQ ID NO:80)

**Toxin Sequence:**

35 Xaa2-Gln-Cys-Cys-Xaa3-Xaa3-Val-Ala-Cys-Asn-Met-Gly-Cys-Xaa1-Xaa3-Cys-Cys-# (SEQ  
ID NO:81)

Name: Da3.3

Species: dalli

40 Cloned: Yes

**DNA Sequence:**

CAAGAAGGGATCGATAGCAGTTCATGATGTCATAAAGCTGGAGCCTGTTGATCATATG  
TCTATTCTGTTCCCTAACTGCTGTTCACTGGAGATCAGCCTGCAGACCAA  
45 TCTGCAGAGCGTATGCAGGACAAAATTCTGAACATCATCCCTTTGATCCC  
GTCAAACGTTGCAACGCGGGGTTTGGCGCTTCGGATGCACGCCTGTTGG

TGACCAAGCTTGTATCGCGGCCTCATCAAGTGTCTAATGAATAAGTAAAATGATTG  
CAGT (SEQ ID NO:82)

**Translation:**

5 MMSKLGVLLIICLFLFPLTAVQLNGDQPADQSAERMQDKISSEHPFFDPVKRCCNAGF  
CRFGCTPCCW (SEQ ID NO:83)

**Toxin Sequence:**

10 Cys-Cys-Asn-Ala-Gly-Phe-Cys-Arg-Phe-Gly-Cys-Thr-Xaa3-Cys-Cys-Xaa4-^ (SEQ ID NO:84)

15 Name: Di3.1  
Species: *distans*  
Cloned: Yes

**DNA Sequence:**

CAAGAGGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCCTGCTGACCATCTT  
TCTGCTTCTGTTCCCCTACTGCTGTTCCGCTGGATGGAGATCAACCCGCAGACGG  
ACTTGCAGAGCGCATGCAGGACGACAGTTCAGCTGCACTGATTAGAGACTGGCTTC  
20 TTCAAACCCGACAGTGTGTGCATCCATGCCATGCACGCCCTGCTGTAGATGAC  
CAGCTTGTATCGCGGCTACGTCAAGTATCTAATGAATAAGTAAGTAAAACGATTG  
CAGT (SEQ ID NO:85)

**Translation:**

25 MMSKLGVLLTIFLLLFPPLTAVPLGDQPADGLAERMQDDSSAALIRDWLLQTRQCCVH  
PCPCTPCCR (SEQ ID NO:86)

**Toxin Sequence:**

30 Xaa2-Cys-Cys-Val-His-Xaa3-Cys-Xaa3-Cys-Thr-Xaa3-Cys-Cys-Arg-^ (SEQ ID NO:87)

35 Name: E3.1  
Species: *ermineus*  
Cloned: Yes

**DNA Sequence:**

ACCTCAAGAGGGATCGATCGCAGTTCATGATGTCTAAACTGGGAGCCTTGTGACC  
ATCTGCTGCTTCTGTTCCCATTACTGCTCTCTGATGGATGGAGATCAGCCTGCAG  
ACCGACCTGCAGAGCGTACGGAGGATGACATTCTGACTACATTCCCTGTTGCA  
40 GTTGGCCATGCCCGATACTCCAACGGTAAACTTGTGTTGTTGCTGCTGGATG  
ATAATGTGTTGATGACCAACTTGTATCACGGCTACGTCAAGTGTCTACTGAATAA  
GTAAAATGATTGCAGTA (SEQ ID NO:88)

**Translation:**

45 MMSKLGALLTICLLFPITALLMDGDQPADRPAERTEDDISSDYIPCCSWPCPRYSNGKL  
VCFCLG (SEQ ID NO:89)

**Toxin Sequence:**

Cys-Cys-Ser-Xaa4-Xaa3-Cys-Xaa3-Arg-Xaa5-Ser-Asn-Gly-Lys-Leu-Val-Cys-Phe-Cys-Cys-Leu-# (SEQ ID NO:90)

5

**Name:** Ge3.2  
**Species:** *generalis*  
**Cloned:** Yes

10 **DNA Sequence:**

GGATCCATGATGTCTAAACTGGGAGTCTTGTGACCACATCTGCTGGTTCTGTTCCCC  
TTACTGCTCTTCACTGGATGGAGAACAAACCTGTAGACCGACATGCCGAGCATATGC  
AGGATGACAATTCTAGCTGCACAGAACCCCTGGGTATTGCCATCAGACAGTGTG  
ACGTTCTGCAACTTGGATGCCAACCTTGTGCCTCACCTGATAACGTGTTGATGAC  
15 CAACTTCTCGAG (SEQ ID NO:91)

**Translation:**

GSMMMSKLGVL LTICLVLFPLTALPLDGEQPVDRHAEHMQDDNSAAQNPWVIAIRQCCT  
FCNFGCQPCCLT (SEQ ID NO:92)

20

**Toxin Sequence:**

Xaa2-Cys-Cys-Thr-Phe-Cys-Asn-Phe-Gly-Cys-Gln-Xaa3-Cys-Cys-Leu-Thr-^ (SEQ ID NO:93)

25

**Name:** Ge3.3  
**Species:** *generalis*  
**Cloned:** Yes

**DNA Sequence:**

GGATCCATGATGTCTAAACTGGGAGTCTTGTGACCACATCTGCTGGTTCTGTTCCCC  
TTACTGCTCTTCACTGGATGGAGAACAAACCTGTAGACCGACATGCCGAGCATATGC  
AGGATGACAATTCTAGCTGCACAGAACCCCTGGGTATTGCCATCAGACAGTGTG  
ACGTTCTGCAACTTGGATGCCAGCCTTGTGCCTCCCTGATAACGTGTTGATGAC  
15 CAACTTCTCGAG (SEQ ID NO:94)

35

**Translation:**

GSMMMSKLGVL LTICLVLFPLTALPLDGEQPVDRHAEHMQDDNSAAQNPWVIAIRQCCT  
FCNFGCQPCCVP (SEQ ID NO:95)

40

**Toxin Sequence:**

Xaa2-Cys-Cys-Thr-Phe-Cys-Asn-Phe-Gly-Cys-Gln-Xaa3-Cys-Cys-Val-Xaa3-^ (SEQ ID NO:96)

45

**Name:**  $\mu$ -GIIA  
**Species:** *geographus*  
**Cloned:** Yes

**DNA Sequence:**

5 GTCGACTCTAGAGGATCCGACAACAAAGAGTCAACCCCCACTGCCACGTCAAGAGCG  
 AAGGCCACAGCTAACGACAAGAGGGATCGATAGCAGTTCATGATGTCTAAACTGGG  
 10 AGTCTTGTGACCATCTGCTGCTTCTGTTCCCTTACTGCTCTCCGATGGATGGA  
 GATGAACCTGCAAACCGACCTGTCGAGCGTATGCAGGACAACATTCTATCTGAGCA  
 GTATCCCTGTTGAGAAGAGAGGAGATTGTCAGTCCGCCAGAAATGCAAAG  
 ACCGACAATGCAAACCCCCAGAGAGATGTCGCTGGACGATAACGTGTTGATGACCAA  
 CTTTATCACGGCTACGTCAGTGTAGTGAATAAGTAAATGATTGAGTCTTGCT  
 15 CAGATTGCTTTGTGTTGGTCTAAAGATCAATGACCAAAACCGTTGTTGATGCG  
 GATTGTCATATATTCTGATTCCAATCCAACACTAGATGATTAATCACGATAGAT  
 TAATTCTATCAATGCCCTGATTTCGTCATATCAGTTGTTATATTATT  
 TTTCTGTCACTGTCTACACAAACGCATGCATGCACGCACACACGCACGC  
 ACGCTGCACAAACATGCGCGCAGCACACACACACACACACACACAAACACA  
 20 15 CACACAAGCAATCACACAATTATTGACATTATTATTATTATTGATGTATTGTTA  
 TTCGTTGCTTGTAGAATAGTTGAGGCCGTTGGATTATTGAACCTGC  
 TTTATTGTATACGAGTACTCGTCTTGAAACACTGCTGAAAATAAAACAAACACT  
 GACGTAGC (SEQ ID NO:97)

**20 Translation:**

MMSKLVLLTICLLLFPLTALPMGDPEANPVERMQDNISSEQYPLFEKRRDCCTPPK  
 KCKDRQCKPQRCCAGR (SEQ ID NO:98)

**Toxin Sequence:**

25 Arg-Asp-Cys-Cys-Thr-Xaa3-Xaa3-Lys-Lys-Cys-Lys-Asp-Arg-Gln-Cys-Lys-Xaa3-Gln-Arg-  
 Cys-Cys-Ala-# (SEQ ID NO:99)

30 **Name:**  $\mu$ -GIIB

**Species:** geographus

**Isolated:** Yes

**Cloned:** Yes

**DNA Sequence:**

35 GGCCAGACGACAACAAAGAGTCAACCCCCACTGCCACGTCAAGAGCGAAGCGCAC  
 AGCTAAGACAAGAGGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGT  
 GACCATCTGCTGCTTCTGTTCCCTTACTGCTCTCCGATGGATGGAGATGAAACCT  
 GCAAACCGACCTGTCGAGCGTATGCAGGACAACATTCTATCTGAGCAGTATCCCTG  
 TTTGAGAAGAGACGAGATTGTCAGTCCGCCAGGAAATGCAAAGACCGACGATG  
 40 45 CAAACCCATGAAATGTCGCTGGACGATAACGTGTTGATGACCAACTTATCAG  
 GCTAGCTCAGTGTAGTGAATAAGTAAATGATTGAGTCTGCTCAGATTGCTT  
 TGTGTTTGGCTAAGATCAATGACCAAAACCGTTGTTGATGCGGATTGTCATATA  
 TTTCTGATTCCAATCCAACACTAGATGATTAAATCACGATAGATTAATTCTATCA  
 ATGCCITGATTTCGTCATATCAGTTGTTATTTATTGTCAGTCTGCTCAGATTGCTT  
 CTACACAAACGCATGCATGCACGCATGCACGCACACACGCACGCACGCTCGCACAA  
 ACATGCGCGCAGCACACACACACACACACACACACACACACACACAGAAGCAATC  
 ACACAATTAGTTGACATTATTATTATTGATGTATTGTTATTGCTTGT

TTTTAGAATAGTTGAGGCCGTCTTTGGATTTATTGAAC TGCTTATTGTATACG  
AGTACTTCGTGCTTGAAACACTGCTGAAATAAAACAAACACTGACGTAGCAAAA  
AAAAAAA (SEQ ID NO:100)

5 **Translation:**

MMSKLGVLLTICLLLPLTALPMGDDEPANRPVERMQDNISSEQYPLFEKRRDCCTPPR  
KCKDRRCKPMKCCAGR (SEQ ID NO:101)

**Toxin Sequence:**

10 Arg-Asp-Cys-Cys-Thr-Xaa3-Xaa3-Arg-Lys-Cys-Lys-Asp-Arg-Arg-Cys-Lys-Xaa3-Met-Lys-  
Cys-Cys-Ala-# (SEQ ID NO:102)

Name:  $\mu$ -GIIC

15 Species: geographus

Isolated: Yes

**Toxin Sequence:**

20 Arg-Asp-Cys-Cys-Thr-Xaa3-Xaa3-Lys-Lys-Cys-Lys-Asp-Arg-Arg-Cys-Lys-Xaa3-Leu-Lys-  
Cys-Cys-Ala-# (SEQ ID NO:103)

Name: Gm3.1

25 Species: gloriamaris

Cloned: Yes

**DNA Sequence:**

CTCACTATAGGAATTCGAGCTCGGTACACGGGATCGATAGCAGTTCATGATGTCTAA  
30 ACTGGGAGCCTTGTGACCATCTGTCTACTTCTGTTCCCTAACTGCTGTTCCGCTG  
GATGGAGATCAACATGCAGACCAACCTGCAGAGCGTCTGCATGACCGCCTCCAAC  
TGAAAATCATCCCTTATATGATCCCCTCAAACGGTGTGCGATGATTGGAAATGCGA  
CTATTCTGCTGGCCTGCTGTATGTTGGATAACCTTGTATCGCGGCCTCGATAA  
GTGTCTAATGAATAAGTAAAACGATTGCAGTAGGC (SEQ ID NO:104)

35

**Translation:**

MMSKLGALLTICLLLFSLTAVPLDGDQHADQPAERLHDRLPTENHPLYDPVKRCCDDSE  
CDYSCWPCCMFG (SEQ ID NO:105)

40 **Toxin Sequence:**

Cys-Cys-Asp-Asp-Ser-Xaa1-Cys-Asp-Xaa5-Ser-Cys-Xaa4-Xaa3-Cys-Cys-Met-Phe-# (SEQ ID  
NO:106)

45 Name: Gm3.2

Species: gloriamaris

Cloned: Yes

**DNA Sequence:**

5 GITCATGATGTCCTAAACTGGGAGTCTTGTGATCATCTGTCTACTTCTGTTCCCTT  
 ACTGCTGTTCCGCTGGATGGAGATCAACCTGCAGACCGATATGCAGAGCGTATGCA  
 GGACGACATTTCATCTGAACATCATCCCATGTTGATGCCGTCAAGAGGGTGTGCCA  
 TCTGTTGGCATGCCGCTCGGATGCTCGCCTGTTGGTGTAGCTAGCTTGTATCG  
 CGGCCTCATCAAGTGAATGCAAA (SEQ ID NO:107)

**Translation:**

10 MMSKLGVLIIICLLLPLTAVPLDGDQPADRYAERMQDDISSEHHPMFDARVGCCHLA  
 CRFGCSPCCW (SEQ ID NO:108)

**Toxin Sequence:**

15 Gly-Cys-Cys-His-Leu-Leu-Ala-Cys-Arg-Phe-Gly-Cys-Ser-Xaa3-Cys-Cys-Xaa4-^ (SEQ ID  
 NO:109)

20 **Name:** Gm3.3  
**Species:** gloriamaris  
**Cloned:** Yes

**DNA Sequence:**

25 GAGACGACAAGGAACAGTCACCCCCACAGCCACGCCAAGAGCAGACAGCCACAGC  
 TACGTGAAGAAGGGTGGAGAGAGGGITCGTGTGAAAATGGGAGTGGTGCTATT  
 CATCTCCTGGTACTGTTCCCTGGCAACGCTCCAGCTGGATGCAGATCAACCTGT  
 AGAACGATATGCGGAGAACAAACAGCTCCTCAACCCAGATGAAAGGAGGGAAATC  
 ATATTGCATGCTCTGGGGACGCGATGCTGTTGGGATGTGTGCGACCACCCGAGT  
 TGTACTTGCTGCGCGGGTAGCGCCAACATCCATGGCGCTGTGCTGGCGGTTTA  
 30 TCCAACAAACGACAGCGTTGTTGATTTCATGTATCATTGCGCCCACGTCTTGTCTA  
 AGAATGACGAACATGATTGCACTCTGGTCAGATTGCTGTTCTTGTACAATAA  
 ATGACAAAACCTCCAAAAAA (SEQ ID NO:110)

**Translation:**

35 MLKMGVVLFIFLVLFPLATLQLDADQPVERYAENKQLNPDERREILHALGTRCCSWD  
 VCDHPSCTCCGG (SEQ ID NO:111)

**Toxin Sequence:**

Cys-Cys-Ser-Xaa4-Asp-Val-Cys-Asp-His-Xaa3-Ser-Cys-Thr-Cys-Cys-Gly-# (SEQ ID NO:112)

40 **Name:** La3.1  
**Species:** laterculatus  
**Cloned:** Yes

45 **DNA Sequence:**  
 CGACCTCAAGAAGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTGA  
 CCATCTGTCTGCTTCTGTTCCCTACTGCTCTCCGATGGATGGAGATCAACCTGC

AGACCGACCTGCAGAGCGTATGCAGGACGTTCATCTGAACAGCATCCCTGTATG  
ATCCCGTCAAACGGTGTGCGACTGGCCATGCAGCGGATGCATCCCTGTTGCTAAT  
AGTAACAACGTGTTGATAACCAACTTCTTACACCAGACTACGTCAAGTGTCTAATGA  
ATAAGTAAAATGATTGCAGT (SEQ ID NO:113)

5

**Translation:**

MMSKLGVLLTICLLLFPPLTALPMGDQPADRPAERMQDVSSSEQHPLYDPVKRCCDWPC  
SGCIPCC (SEQ ID NO:114)

10

**Toxin Sequence:**

Cys-Cys-Asp-Xaa4-Xaa3-Cys-Ser-Gly-Cys-Ile-Xaa3-Cys-Cys-^ (SEQ ID NO:115)

15

Name: La3.2

Species: laterculatus

Cloned: Yes

**DNA Sequence:**

CGACCTCAAGAAGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCITGTTGA  
20 CCATCTGCTGCTTCTGTTCCCTACTGCTCTGGATGGAGATCAACCTGCAGACC  
GAATTGCAGAGCGTATGCAGGACGACATTCTGAGCAGCAGCATCCCTTGAAAAG  
AGACGAGACTGTTGCACACCTCCGAAGAAATGCAGAGACCGACAATGCAAACCTGC  
ACGTTGTTGCGAGGATAACGTGTTGATGACCAACTTGTATCACGGCTACGTCAA  
GTGTCTAGTGAATAAGTAAAACGATTGCAGT (SEQ ID NO:116)

25

**Translation:**

MMSKLGVLLTICLLLFPPLTALDGDPADRLAERMQDDISSEQHPFEKRRDCCTPPKKCR  
DRQCKPARCCGG (SEQ ID NO:117)

30

**Toxin Sequence:**

Arg-Asp-Cys-Cys-Thr-Xaa3-Xaa3-Lys-Lys-Cys-Arg-Asp-Arg-Gln-Cys-Lys-Xaa3-Ala-Arg-  
Cys-Cys-Gly-# (SEQ ID NO:118)

35

Name: La3.3

Species: laterculatus

Cloned: Yes

**DNA Sequence:**

40 GGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCITGTTGACCATCTGTCTGC  
TTCTGTTCCCTTACTGCTCTTCCGATGGATGGAGATCAACTTGCACGCCGATCTGC  
AGAGCGTATGCAGGACAAACATTCTGAGCAGCAGCATCACCTCTTGAAAAGAGAC  
GACCACCATGTTGCACCTATGACGGGAGTTGCCTAAAAGAATCATGCATGCGTAAA  
GCTTGTGCGGATGATAACGTGTTGATGACCAACTTGTATCACGGCTACTCAAGT  
45 GTCTAATGAATAAGTAAAATGATTGCAGTA (SEQ ID NO:119)

**Translation:**

MMSKLGVLLTICLLLFPPLTALPMGDQLARRSAERMQDNISSEQHHLFEKRRPPCCTYD  
GSCLKESCMRKACCG (SEQ ID NO:120)

**Toxin Sequence:**

5 Arg-Xaa3-Xaa3-Cys-Cys-Thr-Xaa5-Asp-Gly-Ser-Cys-Leu-Lys-Xaa1-Ser-Cys-Met-Arg-Lys-  
Ala-Cys-Cys-# (SEQ ID NO:121)

10 Name: La3.3A  
Species: laterculatus  
Cloned: Yes

**DNA Sequence:**

15 GGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTGACCACCTGTC  
TTCTGTTCCCTTACTGCTCTCCGATGGATGGAGATCAACTTGACGCCGACCTG  
CAGAGCGTATGCAGGACAACATTCTATGAGCAGCATCCCTCTTGAAAGGAGA  
CGACCACCATTGTCACCTATGACGGGAGTTGCCTAAAAGAATCATGCAAGCGTAA  
AGCTTGTGCGATAATAACGTGTTGATGACCAACTTGTATCACGGCTACTCAAG  
20 TGTCTAATGAATAAGTAAAATGATTGCAGTA (SEQ ID NO:122)

20 **Translation:**

MMSKLGVLTTICLLLFPPLTALPMGDQLARRPAERMQDNISSEQHPFFERRPPCCTYD  
GSCLKESCKRKACCG (SEQ ID NO:123)

25 **Toxin Sequence:**

Arg-Xaa3-Xaa3-Cys-Cys-Thr-Xaa5-Asp-Gly-Ser-Cys-Leu-Lys-Xaa1-Ser-Cys-Lys-Arg-Lys-  
Ala-Cys-Cys-# (SEQ ID NO:124)

30 Name: Lp3.1  
Species: leopardus  
Cloned: Yes

**DNA Sequence:**

35 GGATCCATGATGTCTAAACTGGGAGTCTTGTGACCGTCTGCTGCTCTGTTCCCC  
TTACTGCTCTCGGCTGGTGGAGATCAACCTGCAGAGCGACCTGCAAAGCGTACGC  
AGGACGACATTCCAGATGGACAGCAGCATCCGTTAAATGATAGGCAGATAAACTGTG  
CCGTGGCCATGCCCTAGTACATGCCGCATCAATGCTGCCATTAAATGATAACGTGTT  
40 GATGACCAACTTCTCGAG (SEQ ID NO:125)

40 **Translation:**

GSMMMSKLGVLTVCLLFPLTALRLVGDQPAERPAKRTQDDIPDGQHPLNDRQINCCP  
WPCPSTCRHQCC (SEQ ID NO:126)

45 **Toxin Sequence:**

Xaa2-Ile-Asn-Cys-Cys-Xaa3-Xaa4-Xaa3-Cys-Xaa3-Ser-Thr-Cys-Arg-His-Gln-Cys-Cys-His-  
(SEQ ID NO:127)

5      Name:      Lv3.1  
Species:      lividus  
Cloned:      Yes

DNA Sequence:  
GGATCCATGATGTCTAAACTGGGAGTCTTGTGACCGTCTGCTGCTCTGTTCCCC  
TTACTGCTCTCGGCTGGTTAGAGATCAACCTGCAGAGCGACCTGCAAAGCGTACGC  
10     AGGACGACATTCCAATGGACAGGATCCGTTAATTGATAGGCAGATAAATTGTTGC  
CCTTGGCCATGCCCTGATTCAATGCCACTATCAATGCTGCCACTGATAACGTGTTGAT  
GACCAACTTCTCGAG (SEQ ID NO:128)

15     Translation:  
GSMMMSKLGVLTVCLLLFPLTALRLVRDQPAERPAKRTQDDIPNGQDPLIDRQINCCPW  
PCPDSCHYQCCH (SEQ ID NO:129)

20     Toxin Sequence:  
Xaa2-Ile-Asn-Cys-Xaa3-Xaa4-Xaa3-Cys-Xaa3-Asp-Ser-Cys-His-Xaa5-Gln-Cys-Cys-His-<sup>^</sup>  
(SEQ ID NO:130)

25     Name:      L3.1  
Species:      lynceus  
Cloned:      Yes

DNA Sequence:  
AAGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTGACCATCTGCTG  
30     CTTCTGTTCCCTTACTGCTCTCCGATGGAGATCAATCTGCAGACCGACTTG  
CAGAGCGTATGCAGGACAACATTCTGAGCAGCATCCCTCTTGAAAAGAGA  
GGACGAGACTGTTGCACACCTCCGAGGAAATGCAGAGACCGAGCCTGCAAACCTCA  
ACGTTGTTGCGGAGGATAAGCTGTTGATGACCAACTTGTATACGGC (SEQ ID  
NO:131)

35     Translation:  
MMSKLGVLTLICLLFPLTALPMGDQSAADRLAERMQDNISSEQHPFFKGRGRDCCTPP  
RKCRDRACKPQRCCGG (SEQ ID NO:132)

40     Toxin Sequence:  
Gly-Arg-Asp-Cys-Cys-Thr-Xaa3-Xaa3-Arg-Lys-Cys-Arg-Asp-Arg-Ala-Cys-Lys-Xaa3-Gln-  
Arg-Cys-Cys-Gly-# (SEQ ID NO:133)

45     Name:      M3.1  
Species:      magus  
Cloned:      Yes

**DNA Sequence:**

CAAGAGGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTGACCATCTG  
 TCTGCTTCTGTTCCCTTAAGTGCCTTCCGATGGATGGAGATGAACCTGCAAACCG  
 ACCTGTCGAGCGTATGCAGGACAACATTCTAGTGCAGTATCCCTGTTGAGAA  
 5 GAGACGAGATTGTCGACTCCGCCAAGAAAATGCAAAGACCGACAATGCAAACCC  
 AGAGATGTTGCGCTGGACGATAACGTGTTGATGACCAACTTATCACGGCTACGTCA  
 AGTGTITAGTGAATAAGTAAATGATTGAGCTTGCAGATTGCTTGTGTTT  
 GGTCTAAAGATCAATGACCAACCGTTGTTGATGCGGATTGTATATATTCTCG  
 ATTCCAATCCAACACTAGATGATTAATCACGATAGATTAATTCTATCAATGCCT  
 10 TGATTTTCGTCTGTCATATCAGTTGTTATTTATTTTCGTCACTGTCTACAC  
 AAACGCATGCATGCACGCATGCACGCACACACGCACGCACGCTCGCACAAACATGC  
 GCGCGCACGCACACACACACACACACACACACACACAGAAGCAATCACAC  
 AATTAGTTGACATTATTATTATTGATGTATTGTTATCGTTGCTTGTGTTT  
 AGAATAGTTGAGGCCGCTTTGGATTATTGAACTGCTTATTGATACGAGTA  
 15 CTTCTGCGGGGAAACACTGCTGAAAAAACAAACACTGACGTAGCAAAAAAA  
 AAAAAA (SEQ ID NO:134)

**Translation:**

MMSKLGVLITICLLLFPALPMDGDEPANRPVERMQDNISSEQYPLFEKRRDCCTPPK  
 20 KCKDRQCKPQRCCAGR (SEQ ID NO:135)

**Toxin Sequence:**

Arg-Asp-Cys-Cys-Thr-Xaa3-Xaa3-Lys-Lys-Cys-Lys-Asp-Arg-Gln-Cys-Lys-Xaa3-Gln-Arg-  
 Cys-Cys-Ala-# (SEQ ID NO:136)

25

Name: M3.2  
 Species: magus  
 Cloned: Yes

30

**DNA Sequence:**

CAAGAGGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTGACCATCTG  
 TCTGCTTCTGTTCCCTTAAGTGCCTTCCGATGGATGGAGATCAACCTGCAAACCG  
 ACCTGTCGAGATCGTATGCAGGACGACATTCTAGTGCAGTATCCCTGTTGATAT  
 35 GAGAAAAAGGTGTTGCCGCCCCGGCGTTCATGCCCGTATATTGAGAGACAATT  
 TTATTGTTGTTGTTAAATGACAACGTGTCGATGACCAACTTCATTATCACGAC  
 TACGCCAAGTGTCTAATGAATAAAATGATTGAGCTCGCTCAGATTGCTT  
 TGTATTGGTCTAAAGATCAATGACCAACCGTTGTTGGATTTCATATA  
 40 TTTCTGAGTCCTATCCAACACTAGATGATTAATCACGATAGATCTGATTTTAT  
 CAAAGGCTGGTTTCGTCTGTCACATCAGTTGTTATTTAATTTCGTCACT  
 GATTACACACACGCATGAACGCACAGAGTACTAACACACACACACACACACA  
 CA  
 CCATCTAGTAGCGCCGCGACGACACACAC (SEQ ID NO:137)

45

**Translation:**

MMSKLGVLITICLLLFPALPMDGDPADQPADRMQDDISSEQYPLFDMRKRCGPG  
 GSCPVYFRDNFICGCC (SEQ ID NO:138)

### Toxin Sequence:

Cys-Cys-Gly-Xaa3-Gly-Gly-Ser-Cys-Xaa3-Val-Xaa5-Phe-Arg-Asp-Asn-Phe-Ile-Cys-Gly-Cys-Cys-^ (SEO ID NO:139)

5

Name: M3.3

Species: magus

Cloned: Yes

10

### DNA Sequence:

### Translation:

25

MMSKLGVLTTICLLLFPLTALPRGDQSVDRAERMQDDISSELHPLSIRKRMCCGESAP  
CPSYFRNSOICHCC (SBO ID NO:141)

### Toxin Sequence:

Met-Cys-Cys-Gly-Xaa<sub>1</sub>-Ser-Ala-Xaa<sub>3</sub>-Cys-Xaa<sub>3</sub>-Ser-Xaa<sub>5</sub>-Phe-Arg-Asn-Ser-Gln-Ile-Cys-His-Cys-Cys-^ (SEO ID NO:142)

30

Name: M3.4

Species: magus

35

### DNA Sequence:

CAAGAGGGATCGATAGCAGTTCATGATGCTAAACTGGGAGTCTTGTGACCATCTG  
TCTGCTCTGTTCCCTTACTGCTCTCCAATGGATGGAGATCAACCTGCAGACCA  
ACCTGCAGATCGTATGCAGGACGACATTCTGAGCAGTATCCCTGTTGATAA  
GAGACAAAAGTGTGCGGCCCCGGCGGTATGCCCCGTATATTACAGACAAATT  
TATTGTGGTTGTGTTAAATGACAACGTGTCGATGACCAACTTCATTATCAGACT  
ACGCCAAGTGTCTAATGAATAAATAAAATGATTGCACTCGCTCAGATTGCTTTT  
GTATTGGCTAAAGATCAATGACCAAACCGTTGTTGGTGGATTTCATATA  
TTCTCGATTCCATCCAACACTAGATGATTAAATCACGATAGATCTGATTTCAT  
CAATGCCTTAATTGGTGTCTGTCATATCAGTTGTTATAT (SEQ ID NO:143)

**Translation:**

MMSKLGVLLTICLLLPLTALPMGDQPADQPADRMQDDISSEQYPLFDKRQKCCPG  
GSCPVYFTDNFICGCC (SEQ ID NO:144)

5 **Toxin Sequence:**

Xaa2-Lys-Cys-Cys-Gly-Xaa3-Gly-Gly-Ser-Cys-Xaa3-Val-Xaa5-Phe-Thr-Asp-Asn-Phe-Ile-Cys-  
Gly-Cys-Cys-^ (SEQ ID NO:145)

10 **Name:** M3.5**Species:** magus**Cloned:** Yes**DNA Sequence:**

15 CAAGAGGGATCGATAGCAGTTCATGATGTCATAACTGGGAGTCTTGTGACCATCTG  
TCTGCCTCTGTTCCCTTAAGCTCTTCCAAATGGATGGAGATCAACCTGCAGACCA  
ACCTGCAGATCGTATGCAGGACGACATTTCATCTGAGCAGTATCCCTGTTGATAA  
GAGACAAAAGTGTGCGGCCCGCGGTTCAATGCCCGTATATTTCAGAGACAATT  
TATTGTGGTTGTGTTAAATGACAACGTGTCGATGACCATCTCATTATCACGACT  
20 ACGCCAAGTGTCTAATGAATAAATAAAATGATTGAGTCAGTCTCGCTCAGATTGCTTT  
GTATTGGTCTAAAGATCAATGACCAACCGTTGTTGGATTTCATATAT  
TTCTCGATTCCATCCAACACTAGATGATTAAATCACGATAGATCTGATTTTT (SEQ  
ID NO:146)

25 **Translation:**

MMSKLGVLLTICLLLPLTALPMGDQPADQPADRMQDDISSEQYPLFDKRQKCCPG  
GSCPVYFRDNFICGCC (SEQ ID NO:147)

**Toxin Sequence:**

30 Xaa2-Lys-Cys-Cys-Gly-Xaa3-Gly-Gly-Ser-Cys-Xaa3-Val-Xaa5-Phe-Arg-Asp-Asn-Phe-Ile-  
Cys-Gly-Cys-Cys-^ (SEQ ID NO:148)

35 **Name:** U001**Species:** magus**Isolated:** No**Toxin Sequence:**

40 Xaa2-Lys-Cys-Cys-Ser-Gly-Gly-Ser-Cys-Xaa3-Leu-Xaa5-Phe-Arg-Asp-Arg-Leu-Ile-Cys-Xaa3-  
Cys-Cys-^ (SEQ ID NO:149)

45 **Name:** Comatose/Death**Species:** marmoreus**Isolated:** Yes**Toxin Sequence:**

Ser-Lys-Gln-Cys-Cys-His-Leu-Ala-Ala-Cys-Arg-Phe-Gly-Cys-Thr-Xaa3-Cys-Cys-Asn-^ (SEQ ID NO:150)

5      Name:      Mr3.1  
Species:      marmoreus  
Cloned:      Yes

**DNA Sequence:**

10      CAAGAAGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTGACCATCTG  
TCTGCTTCTGTTCCCGTTACTGCTCTCCGATGGATGGTGATCAACCTGCAGACCGA  
CTTGTAGAGCGTATGCAGGACAACATTTCATCTGAGCAGCAGTCCCTTCTTGAAAAG  
AGAAGAGGAGGCTGTTGCACACCTCCGAGGAAATGCAAAGACCGAGCCTGCAAAC  
CTGCACGTTGCTGCGGCCAGGATAACGTGTTGATGACCAACTTGTTATCACGGCT  
15      ACGTCAAGTGTCTAGTGAATAAGTAAAACGATTGCAG (SEQ ID NO:151)

**Translation:**

MMSKLGVLLTICLLLFPVTALPMGDQPADRLVERMQDNISSEQHPFFEKRGGCCTPP  
RKCKDRACKPARCCGPG (SEQ ID NO:152)

20      **Toxin Sequence:**  
Arg-Gly-Gly-Cys-Cys-Thr-Xaa3-Xaa3-Arg-Lys-Cys-Lys-Asp-Arg-Ala-Cys-Lys-Xaa3-Ala-Arg-  
Cys-Cys-Gly-Xaa3-# (SEQ ID NO:153)

25      Name:      Mr3.2  
Species:      marmoreus  
Cloned:      Yes

30      **DNA Sequence:**

GAGCTCGGTACCCCGACCTCAAGAGGGATCGATAGCAGTTCATGATGTCTAAACTG  
GGAATCTTGTGACCATCTGTCTACTTCTATTCCCTTACTGCTGTTCCGCTGGATG  
GAGATCAACCTGCAGACCGACCTGCAGAGCGTATGCAGGACGACATTCTGAA  
CATCATCCCTTTTGATCCCGTCAAACGGTGTGCAGGTTATCATGCGGCCTGGGA  
35      TGCCACCCCTGTTGTGGATGACCAGCTTGTATCGCGGCCTCATCAAGTGTCTAAT  
GAATAAGTAAAA (SEQ ID NO:154)

**Translation:**

MMSKLGILLTICLLLFPPLTAVPLGDQPADRPAERMQDDISSEHHPFFDPVKRCCRLSCG  
40      LGCHPCCG (SEQ ID NO:155)

**Toxin Sequence:**

Cys-Cys-Arg-Leu-Ser-Cys-Gly-Leu-Gly-Cys-His-Xaa3-Cys-Cys-# (SEQ ID NO:156)

45      Name:      Mr3.3  
Species:      marmoreus

Cloned: Yes

**DNA Sequence:**

5 GGCCTACACCAAGCTTGCATGCCCTGCAGGTCGACTCTAGAGGGATCCCCGATCGATA  
GCAGTTCATGATGTCTAGACTGGGAGTCTTGTGACCATCTGTCTACTCTGTTCCC  
CTTACTGCTGTCGCTGGATGGAGATCAACCTGCCGACCTGCAGAGCGCCTG  
CAGGACGACATTTCATCTGAACATCATCCCCATTTGATTCCGGCAGAGAGTGTGC  
GGTCGTTCGCATGCCGCTTGGATGCGTGCCTGTGTATGACCAGCTTGTAT  
10 CACGGCCTCATCGAGTGTCTAATGAATAAGTAAAACGATTGCAGTAGGCGGGTACC  
GAGCTCGAATTCC (SEQ ID NO:157)

**Translation:**

MMSRLGVLLTICLLFPLTA VPLDGDQPADRP AERLQDDISSEHHPHFD SGRECCGSFAC  
RFGCVPCCV (SEQ ID NO:158)

15

**Toxin Sequence:**

Xaa1-Cys-Cys-Gly-Ser-Phe-Ala-Cys-Arg-Phe-Gly-Cys-Val-Xaa3-Cys-Cys-Val:^ (SEQ ID  
NO:159)

20

Name: Mr3.4

Species: marmoreus

Cloned: Yes

25

**DNA Sequence:**

CGACCTCAAGAGGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTGA  
CCATCTGTCTACTTCTATTCCCCCTACTGCTGTTCCGCTGGATGGAGACCAACCTGC  
AGACCGACCTGCAGAGCGTATGCAGGACGACATTTCATCTGAACGTCTCCTTTTT  
TGATCGCAGAAACAGTGTGCCATCTGCCGGCATGCCGCTCGGATGTACGCCCTG  
30 TTGTTGGTGTACAGTTGTATCGCGTCCTCATCAAGTGTCTAATGAATAAGTAAA  
ATGATTGCAG (SEQ ID NO:160)

35

**Translation:**

MMSKLGVL LTICLLFPLTA VPLDGDQPADRP AERMQDDISSEHPFFDRSKQCCHLPA  
CRFGCTPCCW (SEQ ID NO:161)

40

Name: Mr3.5

Species: marmoreus

Cloned: Yes

45

**DNA Sequence:**

GGATCCATGATGTCTAAACTGGGAGTCTTGTGACCATCTGTCTGCCCTGTTC

TTACTGCTCTCCGCTGGATGGAGATCAACCTGCAGACCAACGTGCAGAGCGTACG  
CAGGCCGAGAAGCATTCTGCCTGATCCGAGAATGGGCTGTTGCCGTTCCATGC  
AAAACCAGTTGCACTACTTGTGTTGCCGGTGATGATAACGTGTTGATGACCAACTT  
TCTCGAG (SEQ ID NO:163)

5

**Translation:**

GSMSKLGVLLTICLLLPLTALPLGDQPADQRAERTQAEKHSLPDPRMGCCPFPCKT  
SCTTLCG (SEQ ID NO:164)

10

**Toxin Sequence:**

Met-Gly-Cys-Cys-Xaa3-Phe-Xaa3-Cys-Lys-Thr-Ser-Cys-Thr-Thr-Leu-Cys-Cys# (SEQ ID NO:165)

15

**Name:** U014

**Species:** marmoreus

**Isolated:** Yes

**Toxin Sequence:**

20

Cys-Cys-His-Xaa4-Asn-Xaa4-Cys-Asp-His-Leu-Cys-Ser-Cys-Cys-Gly-Ser-^ (SEQ ID NO:166)

**Name:** U017

**Species:** marmoreus

25

**Cloned:** Yes

**DNA Sequence:**

GCCAAGCTTGCATGCCTGCAGGATGACTCTAGAGGATCCCCACCTCAAGAGGGATC  
GATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTGACCATCTGTCTACTTCTGTT  
30 TGCCCTTACTGCTGTTCCGCTGGATGGAGATCAACCTGCAGACCGACCTGCAGAACG  
TATGCAGGACCGACATTCATCTGAACGTCACTCCATGTTGATGCCGTCAAGAGATTG  
TTGCCCGTTGCCGGCATGCCCTTGGATGCAACCCCTGTTGGATGACCAGCTT  
GTTATCGGGACCTCATCAAGTGTCTAATGAATAAGAAAAACGATTGAGTGGGT  
ACCGAGCTCGAATTCC (SEQ ID NO:167)

35

**Translation:**

MMSKLGVLLTICLLLPLTAVPLGDQPADRPAERMQDDISSERHPMFDAVRDCCPLP  
ACPFGCNPCCG (SEQ ID NO:168)

40

**Toxin Sequence:**

Asp-Cys-Cys-Xaa3-Leu-Xaa3-Ala-Cys-Xaa3-Phe-Gly-Cys-Asn-Xaa3-Cys-Cys# (SEQ ID NO:169)

45

**Name:** U019

**Species:** marmoreus

**Isolated:** Yes

**Toxin Sequence:**

Cys-Cys-Ala-Xaa3-Ser-Ala-Cys-Arg-Leu-Gly-Cys-Arg-Xaa3-Cys-Cys-Arg-^ (SEQ ID NO:170)

5

Name: U020  
Species: marmoreus  
Isolated: Yes

10

**Toxin Sequence:**

Cys-Cys-Ala-Xaa3-Ser-Ala-Cys-Arg-Leu-Gly-Cys-Arg-Xaa3-Cys-Cys-Arg-^ (SEQ ID NO:171)

15

Name: U022  
Species: marmoreus  
Isolated: Yes

**Toxin Sequence:**

Cys-Cys-Ala-Xaa3-Ser-Ala-Cys-Arg-Leu-Gly-Cys-Arg-Xaa3-Cys-Cys-Arg-^ (SEQ ID NO:172)

20

Name: U024  
Species: marmoreus  
Isolated: Yes

25

**Toxin Sequence:**

Gly-Cys-Cys-Gly-Ser-Phe-Ala-Cys-Arg-Phe-Gly-Cys-Val-Xaa3-Cys-Cys-Val-^ (SEQ ID NO:173)

30

Name: Nb3.1  
Species: nobilis  
Cloned: Yes

35

**DNA Sequence:**GGATCCATGATGTCTAAACTGGGAGTCTTGTGACCATCTGTCTACTTCTGTTCCCC  
TTACTGCTTCCGCTGGATGAAGATCAACCGGTACACCGACCTGCAGAGCGTATGC  
AGGACATTTCATCTGATCAACATCTCTTGTCTCATCAAACGGTGCTGCGAGT  
TGCCATGCGGGGCCAGGCTTGTGCTCCCTTGTGCTGACATCAATAACGTGTTGATG  
40 ACCAACCTTCTCGAG (SEQ ID NO:174)

45

**Translation:**GSMM\$KLGVLTLTICLLFPLTALPLDEDQPVHRPAERMQDISSDQHLFFDLIKRCCELPC  
GPGFCVPCC (SEQ ID NO:175)**Toxin Sequence:**

Cys-Cys-Xaa1-Leu-Xaa3-Cys-Gly-Xaa3-Gly-Phe-Cys-Val-Xaa3-Cys-Cys-^ (SEQ ID NO:176)

5      Name:      Nb3.2  
Species:      nobilis  
Cloned:      Yes

**DNA Sequence:**

10      GGATCCATGATGTCTAAACTGGGAGTCTTGTGACCATCTGTCTACTTCTGTTCCCC  
TTACTGCTTCCGATGGATGGAGATCAACCTGCAGACCAACCTGCAGATCGTATGC  
AGGACGACATTCATCTGAGCAGTATCCCTGTTGATAAGAGACAAAAGTGTGCA  
CTGGGAAGAAGGGGTATGCTCCGGCAAAGCATGCAAAAATCTCAAATGTTGCTCT  
GGACGATAACGTGTTGATGACCAACTTCTCGAG (SEQ ID NO:177)

**Translation:**

15      GSMMMSKLGVLLTICLLFPLTAFPMDQPADQPADRMQDDISSEQYPLFDKRQKCCT  
GKKGSCSGKACKNLKCCSGR (SEQ ID NO:178)

**Toxin Sequence:**

20      Xaa2-Lys-Cys-Cys-Thr-Gly-Lys-Lys-Gly-Ser-Cys-Ser-Gly-Lys-Ala-Cys-Lys-Asn-Leu-Lys-  
Cys-Cys-Ser-# (SEQ ID NO:179)

25      Name:      Pu3.1  
Species:      pulicarius  
Cloned:      Yes

**DNA Sequence:**

30      GGATCCATGATGTCTAAACTGGGAGTTTGTGACCATCTGTCTGCTTCTGTTCCCC  
TTACTGCTGTTCCGCTGGATGGAGATCAACCTGCAGACCGACCTGCAGAGCGTATGC  
AGGACATTGCAACTGAACAGCAGTCCCTTCTTGATCCCGTCAAACGGTGTGCAACA  
GCTGTTACATGGGATGCATCCCTGTTGCTTAGTAATAACGTGTTGATGACCAAC  
TTTCTCGAG (SEQ ID NO:180)

**Translation:**

35      GSMMMSKLGVLLTICLLFPLTAVPLGDQPADRPAERMQDIATEQHPFFDPVKRCCNSC  
YMGCPCCF (SEQ ID NO:181)

**Toxin Sequence:**

40      Cys-Cys-Asn-Ser-Cys-Xaa5-Met-Gly-Cys-Ile-Xaa3-Cys-Cys-Phe-^ (SEQ ID NO:182)

45      Name:      Qc3.1  
Species:      quercinus  
Cloned:      Yes

**DNA Sequence:**

GGATCCATGATGTCTAAACTGGGAGTCTTGTGACCATCTGTCTGCTTCTGTTCCCC

TTACAGCTCTCAGCTGGATGGAGATCAACCTGCAGACCGACCTGCAGAGCGTACG  
CAGGACATTGCATCTGAACAGTATCGAAAGTTGATCAGAGACAGAGGGTGTGCCA  
GTGGCCATGCCCGGTAGTTGAGATGCTGCCGTACTGGTTAACGTGTGATGACCA  
ACTTTCTCGAG (SEQ ID NO:183)

5

**Translation:**

GSMMMSKLGVL LTICLLL FPLTALQ LDGDQ PADR PAERT QDIASE QYR KFD QR QRCCQW  
PCPGSCRC CRTG (SEQ ID NO:184)

10

**Toxin Sequence:**

Xaa2-Arg-Cys-Cys-Gln-Xaa4-Xaa3-Cys-Xaa3-Gly-Ser-Cys-Arg-Cys-Cys-Arg-Thr-# (SEQ ID  
NO:185)

15

Name: QcIIIA  
Species: quercinus  
Isolated: Yes

**Toxin Sequence:**

20

Cys-Cys-Ser-Gln-Asp-Cys-Leu-Val-Cys-Ile-Xaa3-Cys-Cys-Xaa3-Asn-# (SEQ ID NO:186)

Name: QcIIIB  
Species: quercinus  
Isolated: Yes

**Toxin Sequence:**

Cys-Cys-Ser-Arg-His-Cys-Xaa4-Val-Cys-Ile-Xaa3-Cys-Cys-Xaa3-Asn-? (SEQ ID NO:187)

30

Name: R3.1  
Species: radiatus  
Isolated: Yes  
Cloned: Yes

**DNA Sequence:**

TCAAGAAGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTGACCATCT  
GTCTGCTTCTGTTCCCTTACTGCTCTCCGATGGATGGAGATCAACCTGTAGACCG  
ACTTGAGAGCGTATGCAGGACAACATTCTCATCTGAGCAGCAGCATACCTCTTGA  
GAGACTACCATCGTGTGCTCCCTTAACCTGCGGCTTGCCAGTACCAAGCATGCAA  
ACGTAACCCTGTTGCACAGGATAACGTGTGATGACCAACTTGTTATCACGGCTA  
CGTCAAGTGTCTAGTGAATAAGTAAAACGATTGCAGT (SEQ ID NO:188)

**Translation:**

45

MMSKLGVL LTICLLL FPLTALPMDGDQPVDR LAERM QDNISSEQHTFFEKRLPSCCSLN  
LRLCPVPACKRNPCCTG (SEQ ID NO:189)

**Toxin Sequence:**

Leu-Xaa3-Ser-Cys-Cys-Ser-Leu-Asn-Leu-Arg-Leu-Cys-Xaa3-Val-Xaa3-Ala-Cys-Lys-Arg-Asn-Xaa3-Cys-Cys-Thr-# (SEQ ID NO:190)

5

**Name:** R3.2  
**Species:** radiatus  
**Cloned:** Yes

10 **DNA Sequence:**

AGGTCGACTCTAGAGGATCCCAAGGATCGATAGCAGTTCATGATGTCTAAACTGG  
GAGTCTTGTGACCATCTGTCTGCTCTGTTCCCTACTGCTCTCCGATGGATGG  
AGATCAACCTGCAGACCGACTGCAAGAGCGTATGCAGGACGACATTCTGAGC  
AGCATCCCTTCTTAAAAAGAGACAACAAAGATGTTGCACCGTTAAGAGGATTGT  
15 CCAGTACCAAGCATGCAGAAGTAAACCTTGTGCAAATCATAACGTATTGATGACCA  
ACTTTGTTATCACGGCTACGTCAAGTGTCTAGTGAATAAGTAAAATGATTGCAG  
(SEQ ID NO:191)

**Translation:**20 MMSKLGVLLTICLLLFPPLTALPMGDQPADRLAERMQDDISSEQHPFFKKRQQRCCTV  
KRICPVPACRSKPCCKS (SEQ ID NO:192)**Toxin Sequence:**

25 Xaa2-Gln-Arg-Cys-Cys-Thr-Val-Lys-Arg-Ile-Cys-Xaa3-Val-Xaa3-Ala-Cys-Arg-Ser-Lys-Xaa3-Cys-Cys-Lys-Ser-^ (SEQ ID NO:193)

30 **Name:** R3.3  
**Species:** radiatus  
**Cloned:** Yes

**DNA Sequence:**

ACCTCAAGAAGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTGACC  
ATCTGCTGCTCTGTTCCGTACTGCTCTCCGATGGATGGTATCAACCTGCAG  
35 ACCGACTTGTAGAGCGTATGCAGGACAACATTCTGAGCAGCATCCCTTCTTGG  
AAAAGAGAAGAGGAGGCTGTTGCACACCTCCGAGGAAATGCAAAGACCGAGCCTG  
CAAACCTGCACGTTGCTGCGGCCAGGATAACGTGTTGATGACCAACTTGTATCA  
CGGCTACGTCAAGTGTCTAGTGAATAAGTAAAACGATTGCAGT (SEQ ID NO:194)

40 **Translation:**

MMSKLGVLLTICLLLFPVTALPMGDQPADRLVERMQDNISSEQHPFFEKRGGCCTPP  
RKCKDRACKPARCCGPG (SEQ ID NO:195)

**Toxin Sequence:**

45 Arg-Gly-Gly-Cys-Cys-Thr-Xaa3-Xaa3-Arg-Lys-Cys-Lys-Asp-Arg-Ala-Cys-Lys-Xaa3-Ala-Arg-Cys-Cys-Gly-Xaa3-# (SEQ ID NO:196)

5      **Name:**      Ra3.1  
**Species:**      *rattus*  
**Cloned:**      Yes

**DNA Sequence:**  
GGATCCATGATGTCTAAACTGGGAGTCITGGTGACCATCTGCCTGCTTCTGTTCCCT  
CTTGCTGCTTCCACTGGATGGAGATCAACCTGCAGACCACCCTGCAAAGCGTACG  
10      CAAGATGACAGTCAGCTGCCCTGATCAATGCCTGGCTTGATGAATCCCAGACTTGC  
TGCAGTAACTGCGGTGAAGATTGTGATGGTTGCCAGTAACGTGTTGATGACCAA  
CTTTCTCGAG (SEQ ID NO:197)

15      **Translation:**  
GSMMMSKLGVLVTICLLLFPPLAAFPLGDQPADHPAKRTQDDSSAALINAWLDESQTCCS  
NCGEDCDGCCQ (SEQ ID NO:198)

20      **Toxin Sequence:**  
Xaa2-Thr-Cys-Cys-Ser-Asn-Cys-Gly-Xaa1-Asp-Cys-Asp-Gly-Cys-Cys-Gln-^ (SEQ ID  
NO:199)

25      **Name:**      Sm3.1  
**Species:**      *stercusmuscarum*  
**Cloned:**      Yes

30      **DNA Sequence:**  
GACCTCAAGAGGGATCGATAGCAGTCGTGATGTCTAAACTGGGAGTCITGGTGCAC  
CATCTGTCTGCTTCTGTTCCCTTACTGCTCTCCGATGGATGGAGATCAACCTGCA  
GACCAACCTGCAGATCGTATGCAGGACGACATTTCATCTGAGCAGTATCCCTGTT  
35      GATAAGAGACAAAAGTGTGCACTGGGAAGAAGGGGTATGCTCCGGCAAAGCAT  
GCAAAAATCTCAAATGTTGCTCTGGACGATAACGTGTTGATGACCAACTTGTATC  
ACGGCTACGTCAAGTGTCAATGAATAAGTAAAACGATTGCAGT (SEQ ID NO:200)

35      **Translation:**  
MSKLGVLVTICLLLFPPLALPMGDQPADQPADRMQDDISSEQYPLFDKRQKCCGTGKK  
GSCSGKACKNLKCCSGR (SEQ ID NO:201)

40      **Toxin Sequence:**  
Xaa2-Lys-Cys-Cys-Thr-Gly-Lys-Lys-Gly-Ser-Cys-Ser-Gly-Lys-Ala-Cys-Lys-Asn-Leu-Lys-  
Cys-Cys-Ser-# (SEQ ID NO:202)

45      **Name:**      U034  
**Species:**      *stercusmuscarum*  
**Isolated:**      Yes  
**Cloned:**      Yes

**DNA Sequence:**

GATCGATAGCAGTCGTGATGTCTAAACTGGGAGTCTTGTGACCATCTGCTGCTT  
CTGTTCCCTTACTGCTCTCCGATGGATGGAGATCAACCTGCAGACCAACCTGCA  
5 GATCGTATGCAGAACGACATTTCATCTGAGCAGTATCCCTGTTGATAAGAGACAA  
AAAGTGTGCGGCCCGCGCGTCATGCCCGAGATATTCAAAGACAATTATTTGT  
GGTTGTTGTTAAATGACAACGTGTCGATGACCAACTCGTTATCACGACTTCGCCAA  
GTGTCTAATGAATAAGTAAAACGATTGCAGT (SEQ ID NO:203)

**10 Translation:**

MSKLGVLLTICLLFPLTALPMGDQPADQPADRMQNDISSEQYPLFDKRQKCCGP GAS  
CPRYFKDNFICGCC (SEQ ID NO:204)

**Toxin Sequence:**

15 Xaa2-Lys-Cys-Cys-Gly-Xaa3-Gly-Ala-Ser-Cys-Xaa3-Arg-Xaa5-Phe-Lys-Asp-Asn-Phe-Ile-  
Cys-Gly-Cys-Cys-^ (SEQ ID NO:205)

20 Name: S3.1

Species: striatus

Cloned: Yes

**DNA Sequence:**

25 CGACCTTCAAGAGGGATCGATAGCAGTCGCATGTCTAAACTGGGGTATTGTTG  
ACCATCTGCTGCTCTGTTCCCTTACTGCTCTCCGATGGATGAAGATCAACCTG  
CAGACCAACTTGAAGATCGTATGCAGGACGACATTTCATCTGAGCAGTATCCCTCGT  
TTGTTAGGAGACAAAAGTGTGCGCGAAGGCTCGTCATGCCCAAATATTCAA  
AACAAATTATTTGTGGTTGTTGTTAAATGACAACGTGTCGATGACCAACTCGTTA  
TCACGACTACGCCAAGTGTCTTGTCTAATGATAATAAAATGATGCC (SEQ ID NO:206)

30

**Translation:**

MSKLGVLLTICLLFPLTALPMDEDQPADQLEDRMQDDISSEQYPSFVRRQKCCGEGSS  
CPKYFKNNFICGCC (SEQ ID NO:207)

35

**Toxin Sequence:**

Xaa2-Lys-Cys-Cys-Gly-Xaa1-Gly-Ser-Ser-Cys-Xaa3-Lys-Xaa5-Phe-Lys-Asn-Asn-Phe-Ile-Cys-  
Gly-Cys-Cys-^ (SEQ ID NO:208)

40

Name: S3.2

Species: striatus

Cloned: Yes

**DNA Sequence:**

45

GGATCCATGATGTCTAAACTGGGAGTCTTGTGACCGTCTGCTGCTTCTGTTCCCG  
TTACTGCTCTCCGCTGGATGGAGATCAACCTGCAGACCGACCTGCAGAGCGTATGC  
AGGACGACATTTCATCTGACGAGCATTCCCTGTTGATAAGAGACAAAATGTTGCA

ATGGGGATGCTCCAGCAAATGGTGCAGAGATCACGCACGTGTTGCGGTCGATGA  
TAACGTGTTGATGACCAACTTCTCGAG (SEQ ID NO:209)

**Translation:**

5 GSMMMSKLGVLTVCLLFPLTALPLDGDQPADRPAAERMQDDISSDEHPLFDKRQNCCN  
GGCSSKWC RDHARCCGR (SEQ ID NO:210)

**Toxin Sequence:**

10 Xaa2-Asn-Cys-Cys-Asn-Gly-Gly-Cys-Ser-Ser-Lys-Xaa4-Cys-Arg-Asp-His-Ala-Arg-Cys-Cys #  
(SEQ ID NO:211)

15 Name: Ts3.1  
Species: tessulatus  
Cloned: Yes

**DNA Sequence:**

20 GGATCCATGATGTCTAAACTGGGAGTCTTGTGACCATGTGTGCTCTGTTCCCC  
TTACTGCTGTTCCGCTGGATGGAGATCAACCTGCAGACCGACCTGCAGAGCGTAGG  
CAGGACATTGCAACTGACGATCATCCTTGTGATCCCGTCAAACGGTGTGCCAC  
AAATGCTATATGGGATGCATCCCTTGTGCATTAGTAACGTGTTGATGACCAACTT  
TCTCGAG (SEQ ID NO:212)

**Translation:**

25 GSMMMSKLGVLTMCLLFPLTAVPLDGDQPADRPAAERRQDIATDDHPLFDPVKRCCHK  
CYMGCIPCCI (SEQ ID NO:213)

**Toxin Sequence:**

30 Cys-Cys-His-Lys-Cys-Xaa5-Met-Gly-Cys-Ile-Xaa3-Cys-Cys-Ile-^ (SEQ ID NO:214)

35 Name: Ts3.2  
Species: tessulatus  
Cloned: Yes

**DNA Sequence:**

40 GGATCCATGATGTCTAAACTGGGAGTCTTGTGACCATCTGTGTGCTCTGTTCCCC  
TTACTGCTGTTCCGCTGGATGGAGATCAACCTGCAGACCAACCTGCAGAGCGTAGG  
CAGAACGAGCAGCATCCCTGTATGATCAGAAAAGAAAGTGTGCGGGCCGCATG  
CGCCATGAGCTGCGGCATGGCTAGGTGTGCTATTATGATAACGTGTTGATGACCA  
ACTTCTCGAG (SEQ ID NO:215)

**Translation:**

45 GSMMMSKLGVLTLICVLLFPLTAVPLDGDQPADQPAERTQNEQHPLYDQKRKCCRPPCA  
MSCGMARCCY (SEQ ID NO:216)

**Toxin Sequence:**

Lys-Cys-Cys-Arg-Xaa3-Xaa3-Cys-Ala-Met-Ser-Cys-Gly-Met-Ala-Arg-Cys-Cys-Xaa5-^ (SEQ ID NO:217)

5      Name:      Circling  
 Species:      textile  
 Isolated:      Yes  
 Cloned:      Yes

10     **DNA Sequence:**  
 GAGTCAACCCACTGTCACGCCAAGAGCGGACGCCACAGCTAAGGCAAGAAGGATC  
 GATAGCAGTTCATGATGCTAAACTGGGAGCCTGTTGACCATCTGTCTACTTCTGT  
 TTTCCCTTACTGCTGTTCCGCTGGATGGAGATCAACATGCAGACCAACCTGCACAGC  
 GTCTGCAGGACCGCATTCCAACCTGAAGATCATCCCTTATTGATCCAAACAAACGGT  
 15     GTTGCCTCGCCGGTGGCATGCAACATGGGATGCAAGCCTGTTGATGACCAGCTT  
 TGTTATCGCGGTCTCATGAAGTGTCTAAATGAATAAGTAAACGATTGCAGTTCGTT  
 CAGATTTGCTGTTGTATTTGGTCTAAAGATTAATGACCAAACACTGTTCTTGATCCG  
 GATTTACGTATTCCTGATTCCATTCAACACTAGATAAGTTAATCACGACAGAT  
 20     CTGATTTCCATCAATGCCCTGCTTTGGTCTGTATATAATCTGTTATATTAA  
 TTTCTCGTCACTTCACACACACACACACACACACGCGCGC (SEQ ID  
 NO:218)

25     **Translation:**  
 MMSKLGALLTICLLLFSLTAVPLDGDQHADQPAQRLQDRIPTEDHPLFDPNKRCPPVA  
 CNMGCKPCCG (SEQ ID NO:219).

30     **Toxin Sequence:**  
 Cys-Cys-Xaa3-Xaa3-Val-Ala-Cys-Asn-Met-Gly-Cys-Lys-Xaa3-Cys-Cys-Gly-^ (SEQ ID  
 NO:220)

35     Name:      Scratcher I  
 Species:      textile  
 Cloned:      Yes

40     **DNA Sequence:**  
 GGATCCAGACGACAAAGAAGAGTCACCCACTGCCACGTCAAGAGCAGAGGCCAC  
 AGCTAAGACAAGAAGGATCGATAGCAGTTCATGATGTTAAACTGGGAGTCTTGT  
 GACCATCTGTCCTCTGTTTCCCTTAATGCTGTTCCGTTGGATGGAGATCAACCT  
 45     GCAGACCAACCTGCAGAGCGTCTGCTGGACATTCATTGAAAATAATCCCTT  
 TATGATCCGCCAAACGGTGTGCAGGACTTGCTCGGTGCACACCTGTTGGAT  
 TGACCAGCCTCATCAAGTGTCTAACGAATAAGTAAAGCGATTGCAGTCTCGTTAG  
 ATTTACTTTGTATTCTGGTCTAAAGATTAATGACCAAACCTCTTGTGATCCGGAT  
 GTACATATAATTCTCGATTCCATCCAACGCTAGATAAGCTAATCACGACAGATCTG  
 ATTTCTGTCAATGCCCTGCTTTGGTCTCTCATATCACTCTGTTATATTAAATT  
 CTCGTCACTATATATATACACACACACACACGGAAATTCCGATTGTCCAGTA  
 CCGTTCTGGGATCGAGGTATTGCTGCGATGGCTTATTCTGACTCTTCTGCG

CTTGATAGTGTCTTCTACTCCCACATCTGTGCTACCCCTGGCTTGATCTTGATAGG  
 CGTGTGCCCTTCACTGGTTATAAACCCCTGTATCCTACTCTGGACGCCCTGGGG  
 GCCCAACCTCCAAATAAAGCGACATCCAATGAAAAAA (SEQ ID NO:221)

5 **Translation:**

MMFKLGVLLTICLLLFSLNAVPLDGDQPADQPAERLLDDISFENNPFYDPAKRCRCCRTCF  
 GCTPCCG (SEQ ID NO:222)

**Toxin Sequence:**

10 Cys-Cys-Arg-Thr-Cys-Phe-Gly-Cys-Thr-Xaa3-Cys-Cys-# (SEQ ID NO:223)

Name: Tx3.1

Species: textile

15 Cloned: Yes

**DNA Sequence:**

GGAACAGTCACCCCCACAGCCACGCCAAGAGCAGACAGCCACAGCTACGTGAAGA  
 AGGGTGGAGAGAGGTTCATGATGTTGAAAATGGGAGTGGTGTATTCATCTTCTGG  
 20 TACTGTTCCCTGGCAACGCTCCAGCTGGATGCAGATCAACCTGTAGAACGATATG  
 CGGAGAACAAACAGCTCCTCAACCCAGATGAAAGGAGGGAAATCCTATTGCCTGCT  
 CTGAGGAAGTTCTGCTGTGATTGAAATTGGTGCCACATTCCGGATTGTGAGTGCTGC  
 TACGGTTAGCGCCGAACATCCATGGCACTGTGCTGGCGGGTTCATCCAAACAACG  
 ACAGCGTTGATTGACTCTGGTTGAGATTGATGTATCATTGCGCCCACGTCTTGTCTAAAGAATGACG  
 25 AACATGATTGACTCTGGTTGAGATTGATGTATCATTGCGCCCACGTCTTGTCTAAAGAATGACG  
 TCC (SEQ ID NO:224)

**Translation:**

MMLKMGVVLFIFLVLFPLATLQLDADQPVERYAENKQLNPDERREILLPALRKFCDS  
 30 NWCHISDCECCY (SEQ ID NO:225)

**Toxin Sequence:**

Phe-Cys-Cys-Asp-Ser-Asn-Xaa4-Cys-His-Ile-Ser-Asp-Cys-Xaa1-Cys-Cys-Xaa5-# (SEQ ID  
 NO:226)

35

Name: U031

Species: textile

Isolated: Yes

40 Cloned: Yes

**DNA Sequence:**

CAAGGAACAGTCACCCCCACAGCCACGCCAAGAGCAGACAGCCACAGCTACGTGA  
 AGAAGGGTGGAGAGAGGTTGTGATGTTGAAAATGGGAGTGGTGTATTCATCTTC  
 45 CTGGTACTGTTCCCTGGCAACGCTCCAGCTGGATGCAGATCAACCTGTAGAACGA  
 TATGCGGAGAACAAACAGCTCCTCAGCCCAGATGAAAGGAGGGAAATCATATTGCA  
 TGCTCTGGGGACGCGATGCTGTTGGATGTGCGACCACCGAGTTGTACTTG

CTGCGGTTAGCGCCGAACATCCATGGCGCTGTGCTGGCGGTTATCCAACAACG  
ACAGCGTTGTTGATTTCATGTATCATTGCGCCCACGTCTCTGTCTAAGAATGACG  
AACATGATTGCACTCTGGTTCAAGATTCGTGTTCTTGACAATAATGACAAAAA  
CNCC (SEQ ID NO:227)

5

**Translation:**

MLKMGVVLFIFLVLFPLATLQLDADQPVERYAENKQLLSPDERREIILHALGTRCCSWD  
VCDHPSCTCCG (SEQ ID NO:228)

10

**Toxin Sequence:**

Cys-Cys-Ser-Xaa4-Asp-Val-Cys-Asp-His-Xaa3-Ser-Cys-Thr-Cys-Cys-# (SEQ ID NO:229)

15

Name: U032  
Species: textile  
Isolated: Yes  
Cloned: Yes

**DNA Sequence:**

20

GGATCCATGATGTCTAAACTGGGAGTCTTGTGACCATCTGTCTGCTTCTGTTCCCC  
TTACTGCTCTCCGCTGGATGGAGATCAACCCGCAGACCAAGCTGCAGAGCGTATG  
CAGGCCGAGCAGCATCCCTGTTGATCAGAAAAGACGGTGCTGCAAGTTCCATG  
CCCCGATAGTTGCAGATATTGTGTTGCGGGTGATGATAACGTGTTGATGACCAACT  
TTCTCGAG (SEQ ID NO:230)

25

**Translation:**

GSMMMSKLGVLLLTICLLFPLTALPLDGDQPADQAAERMQAEQHPLFDQKRRCCFKPCP  
DSCRYLCCG (SEQ ID NO:231)

30

**Toxin Sequence:**

Arg-Cys-Cys-Lys-Phe-Xaa3-Cys-Xaa3-Asp-Ser-Cys-Arg-Xaa5-Leu-Cys-Cys-# (SEQ ID NO:232)

35

Name: T3.1  
Species: tulipa  
Cloned: Yes

**DNA Sequence:**

40

CGACCTCAAGAGGGATCGATAGCAGTTCATGTCTAAACTGGGAGTCTTGTGACAA  
TCTGCTGCTCTGTTCCCTTACTGCTCTGCCGATGGATGGAGATGAACCTGCAG  
ACCGACCTGCAGAGCGTATGCAGGACAACATTCTGAGCAGCATCCCTGTTG  
AGGAGAGACACGGATGTTGCAAGGGGCCGAAGGATGCTCCTCCAGAGAATGCAG  
ACCCCAACATTGTTGCGGTGACGATAACGTGTTGAGGGCCAACTTGTTATCACGG  
45 CTACGTCAAGTGTAGTGAATAAGTAAATGATTGCAG (SEQ ID NO:233)

**Translation:**

MSKLGVLLTICLLLFPPLTALPMDGDEPADRPAERMQDNISSEQHPLFEERHGCGCKGPEG  
CSSRECRPQHCCGRR (SEQ ID NO:234)

**Toxin Sequence:**

5 His-Gly-Cys-Cys-Lys-Gly-Xaa3-Xaa1-Gly-Cys-Ser-Ser-Arg-Xaa1-Cys-Arg-Xaa3-Gln-His-  
Cys-Cys-# (SEQ ID NO:235)

10 Name: Fi3.1

Species: *figulinus*

Cloned: Yes

**DNA Sequence:**

15 CAAGAAGGATCGATAGCAGTTCATGATGCTAAACTGGGAGTCCTGCTGACCATCT  
GCTCTGCTTCTGATTCCCTTACTGCTCTTCGCTGGATGGAGATCAACCTGCAGACC  
GACCTGCAGAGCGTATGCAGGATGGAATTTCATCTGAACAGCATCCCATGTTGATC  
CCGTCAGACGGTGTGCCCCATGCAACATAGGATGCGTACCTTGTGTTGAT  
GACCAGTTTGTATCGCGGCCTCATCAAATGTCTAATGAATAAGTAAAACGATTGC  
AGT (SEQ ID NO:236)

20

**Translation:**

MMSKLGVLLTICLLIPLTALSLGDQPADRPAERMQDGISSEQHPMFDPVRCCCPWPC  
NIGCVPCC (SEQ ID NO:237)

25

**Toxin Sequence:**

Cys-Cys-Xaa3-Xaa4-Xaa3-Cys-Asn-Ile-Gly-Cys-Val-Xaa3-Cys-Cys-^ (SEQ ID NO:238)

30 Name: Fi3.2

Species: *figulinus*

Cloned: Yes

**DNA Sequence:**

35 CAAGAGGGATCGATAGCAGTTCATGATGTTAAACTGGGAGTCCTGTTGACCATCTG  
TATGCTTCTGTTCCCTTACTGCTCTCCGCTGGATGGAGAGCAACCTGCAGACCA  
ACCTGCAGAGCGCATGCAGTATGACATGTTACGTGCAATGAATCCCTGGTTGATCC  
CGTCAAAAGGTGCTGCTGAAGAACTGCGCAGTATGCATCCCTGTTGCCCGTAAC  
GACCAGCTGATTATCGCGCCAAGGCTCTAATGAATAAGTAAAACGATTGCAGT  
(SEQ ID NO:239)

40

**Translation:**

MMFKLGVLLTICMLLFPPFTALPLDGEQPADQPAERMQYDMLRAMNPWFDPVKRCCSK  
NCAVCIPCCP (SEQ ID NO:240)

45

**Toxin Sequence:**

Cys-Cys-Ser-Lys-Asn-Cys-Ala-Val-Cys-Ile-Xaa3-Cys-Cys-Xaa3-^ (SEQ ID NO:241)

5      Name:      Fi3.3  
Species:      *figulinus*  
Cloned:      Yes

DNA Sequence:  
CAAGAGGGATCGATAGCAGTTCATGATGTCATAACTGAGAGCTTGTGACCTTATG  
TCTGCTTCTGTTCCCTTACTGCTCTCCGCTGAATGAAGATCAACCTGCAGAGCGT  
10     ATGCAGGACGACAATTCATCTGAGCAGCACCCCTGTATGACCACAAACGAAAGTG  
TTGCCGGTGGCCATGCCCGCAAGATGCGGCTTGTGCTGTAAACGTGTTGG  
CCAACTTGTTATCACGCCACGTCAAATGTTAATGAATAAGTAAAACGATTGCAG  
T (SEQ ID NO:242)

15     Translation:  
MMSKLRVLLTLCLLFPLTALPLNEDQPAERMQDDNSSEQHPLYDHKRKCCRWPCTPAR  
CGSCCL (SEQ ID NO:243)

20     Toxin Sequence:  
Cys-Cys-Arg-Xaa4-Xaa3-Cys-Xaa3-Ala-Arg-Cys-Gly-Ser-Cys-Cys-Leu-^ (SEQ ID NO:244)

25     Name:      Fi3.4  
Species:      *figulinus*  
Cloned:      Yes

DNA Sequence:  
CAAGAGGGATCGATAGCAGTTCATGATGTCATAACTGGGAGCTTGTGACCTTATG  
TCTGCTTCTGTTCCCTGACTGCTCTCCGCTGGATGAAGATCAAGCTGCAGACCG  
30     ACCTGCAGAGCGTATGCAGGGCATGTCATCTGAACAGCATCCCTTCTTGATCCGT  
CAAACGGTGTGCGAGTTGTCACGCTGCCTGGATGCGTCCCTTGTGCACATCTTA  
ATAACGTGTGGATGACCAACTGTGTTATCACGCCACGTCAAAGTGTCTAATGAATA  
AGTAAAATGATTGCAGT (SEQ ID NO:245)

35     Translation:  
MMSKLGVLTLCLLFPLTALPLDEDQAADRPAERMQGMSSEQHPFFDPVKRCCELSR  
CLGCVPCCTS (SEQ ID NO:246)

40     Toxin Sequence:  
Cys-Cys-Xaa1-Leu-Ser-Arg-Cys-Leu-Gly-Cys-Val-Xaa3-Cys-Cys-Thr-Ser-^ (SEQ ID NO:247)

45     Name:      Fi3.5  
Species:      *figulinus*  
Cloned:      Yes

DNA Sequence:

15 CAAGAGGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTGACCTATG  
TCTGCTTCTGTTCCCTGACTGCTCTCCGCTGGATGAAGATCAACCTGCAGACCG  
ACCTGCAGAGCGTATGCAGGGCATGTCATCTGAACAGCATCCCTTGTGATCCCGT  
CAAACGGTGTGCGAGTTGTCAAAATGCCATGGATGCGTCCCTGTGCATAACCTTA  
ATAACGTGCGGATGACCAACTGTGTTATCACGGCCACGTCAAGTGTCTAACCTTA  
AGTAAAATGATTGCAGT (SEQ ID NO:248)

**Translation:**

10 MMSKLGVLTLCLLFPLTALPLDEDQPADRPAAERMQGMSSEQHPFFDPVKRCCELSK  
CHGCVPCCIP (SEQ ID NO:249)

**Toxin Sequence:**

Cys-Cys-Xaa1-Leu-Ser-Lys-Cys-His-Gly-Cys-Val-Xaa3-Cys-Cys-Ile-Xaa3-^ (SEQ ID NO:250)

15 Name: Qc3.2  
Species: quercinus  
Cloned: Yes

20 DNA Sequence:  
CAAGAGGGATCGATAGCAGTTCATGATGTCTAAACTCGGAGTCTTGTGACCATCTG  
TCTGGTTCTGTTCCCTTACAGCTCTCAGCTGGATGGAGATCAACCTGCAGACCG  
ACCTGCAGAGCGTACGCAGGACATTTCATCTGAACAGTATCGAAAGTTGATCAGA  
GACAGAGGTGTGCCGGTGGCCATGCCCGTAGTTGCAGATGCTGCCGTATCGTT  
25 AACGTGTTGGTGACCAGCTTGTATCACGACCACGCCAAGTGTCTAACGAATAAGT  
AAAATGATTGCAGT (SEQ ID NO:251)

**Translation:**

30 MMSKLGVLTLICLVLFPLTALQLDGDQPADRPAAERTQDISSEQYRKFDQRQRCCRWP  
GSCRCCRYSR (SEQ ID NO:252)

**Toxin Sequence:**

Xaa2-Arg-Cys-Cys-Arg-Xaa4-Xaa3-Cys-Xaa3-Gly-Ser-Cys-Arg-Cys-Xaa5-Arg-^  
(SEQ ID NO:253)

35 Name: Qc3.3  
Species: quercinus  
Cloned: Yes

40 DNA Sequence:  
CAAGAGGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTGACCATCTG  
TCTGCTTCTGTTCCCTTACTGCTCTCCACTGGATGGAGATCAACCTGCAGATCAA  
TCTGCAGAGCGACCTGCAGAGCGTACGCAGGACATTCAAGCAGCATCCGTATA  
45 TGATCCGAAAGAAGGTGTGCCGTATCCATGCCCGACAGCTGCCACGGATCTG  
CTGCTATAAGTGATAACATGTTGATGGCCAGCTTGTATCACGGCCACGTCAAGTG  
TCTAATGAATAAGTAAACGATTGCAGT (SEQ ID NO:254)

**Translation:**

MMSKLGVLLTICLLFPLTALPLGDQPADQSAERPAERTQDDIQQHPLYDPKRRCCRY  
PCPDSCHGSCCYK (SEQ ID NO:255)

5

**Toxin Sequence:**

Arg-Cys-Cys-Arg-Xaa5-Xaa3-Cys-Xaa3-Asp-Ser-Cys-His-Gly-Ser-Cys-Cys-Xaa5-Lys-^ (SEQ ID NO:256)

10

**Name:** Wi3.1  
**Species:** *wittigi*  
**Cloned:** Yes

15

**DNA Sequence:**

GGATCCATGATGTCTAAACTGGGAGTCTTGTGACCATCTGCTGCTTCTGTTCCCA  
TTACTGCTCTCCGGTGGGTGGAGATCAGCCTGCAGACCGACTGCAGAGCGTATGC  
AGGACGACACTTCATCTGAGCAGCATCCCTTGAAAAGAGACTACCATCATGTG  
GACTTTGAGAGGCTTGCGTAGTACCAGCATGTCATCAGTGTGCACAGGA  
TAACGTGTTGATGACCAACTTCTCGAG (SEQ ID NO:257)

20

**Translation:**

MMSKLGVLLTICLLFPITALPVGGDQPADRLAERMQDDTSSEQHPFEKRLPSCCDFERL  
CVVPACIRHQCTG (SEQ ID NO:258)

25

**Toxin Sequence:**

Leu-Xaa3-Ser-Cys-Cys-Asp-Phe-Xaa1-Arg-Leu-Cys-Val-Val-Xaa3-Ala-Cys-Ile-Arg-His-Gln-Cys-Cys-Thr-# (SEQ ID NO:259)

30

**Name:** bt3a  
**Species:** *betulinus*  
**Isolated:** Yes

35

**Toxin Sequence:**

Cys-Cys-Lys-Gln-Ser-Cys-Thr-Thr-Cys-Met-Xaa3-Cys-Cys-Xaa4-^ (SEQ ID NO:260)

40

**Name:** T3.2  
**Species:** *tulipa*  
**Cloned:** Yes

45

**DNA Sequence:**

GGATCCATGATGTCTAAACTGGGAGTCTTGTGACAATCTGCTGCTTCTGTTCCCA  
TTACTGCTCTGCCGATGGATGGAGATGAACCTGCAGACCGACCTGCAGAGCGTATG  
CAGGACAAACATTTCATCTGAGCAGCATCCCTGTTGAGGAGAGACACGGATGTTG

CGAGGGGCCGAAGGGATGCTCCTCCAGAGAACATGCAGACCCAACATTGTTGCCGTC  
GACGATAACGTGTTGATGACCAACTNTCTCGAG (SEQ ID NO:261)

**Translation:**

5 MMSKLGVLLTICLLLFPPLTALPMGDPEPADRPAERMQDNISSEQHPLFEERHGCCEGPK  
GCSSRECRPQHCCGRR (SEQ ID NO:262)

**Toxin Sequence:**

10 His-Gly-Cys-Cys-Xaa1-Gly-Xaa3-Lys-Gly-Cys-Ser-Ser-Arg-Xaa1-Cys-Arg-Xaa3-Gln-His-  
Cys-Cys# (SEQ ID NO:263)

Name: A3.5  
Species: aurisiacus  
15 Cloned: Yes

**DNA Sequence:**

20 GGATCCATGATGTCTAAACTGGGAGTCTTGTGACCATCTGTCTACTTCTGTTCCCC  
TTACTGCTTTCCGATGGATGGAGATCAACCTGCAGACCAACCTGCAGATCGTATGC  
AGGACGACATTCATCTGAGCAGTATCCCTGTTGATAAGAGAGACAAAAGTGTGCA  
CTGGGAGGAAGGGGTATGCTCCGGCAAAGCATGCAAAATCTCAAATGTTGCTCT  
GGACGATAACGTGTTGATGACCAACTTCTCGAN (SEQ ID NO:264)

**Translation:**

25 MMSKLGVLLTICLLLFPPLTAFPMGDQPADQPADRMQDDISSEQYPLFDKRQKCCTGR  
KGSCSGKACKNLKCCSGR (SEQ ID NO:265)

**Toxin Sequence:**

30 Xaa2-Lys-Cys-Cys-Thr-Gly-Arg-Lys-Gly-Ser-Cys-Ser-Gly-Lys-Ala-Cys-Lys-Asn-Leu-Lys-  
Cys-Cys-Ser# (SEQ ID NO:266)

Name: Bt3.5  
Species: betulinus  
35 Cloned: Yes

**DNA Sequence:**

40 GGATCCATGATGTCTAAACTGGGAGTCTTGTGACCATCTGTCTGCTTCTGTTCCCC  
TTACTGCTTTCCGTTGGATGGAGATCAACCTGCAGACCAACCTGCAGAGCGTATGC  
AGAACGAGCAGCATCCCTCGTTGATCAGAAAAGAAGGTGCTGCCGGTGGCCATGC  
CCCAGTATATGCGGCATGGCTAGGTGTTGCTTCGTATGATAACGTGTTGATGACCA  
ACTTCTCGAG (SEQ ID NO:267)

**Translation:**

45 MMSKLGVLLTICLLLFPPLTAVPLDGDQPADQPAERMQNEQHPSFDQKRRCCRWPCPSIC  
GMARCCFVMITC (SEQ ID NO:268)

**Toxin Sequence:**

Arg-Cys-Cys-Arg-Xaa4-Xaa3-Cys-Xaa3-Ser-Ile-Cys-Gly-Met-Ala-Arg-Cys-Cys-Phe-Val-Met-Ile-Thr-Cys-^ (SEQ ID NO:269)

5

Name: Bt3.6  
Species: betulinus  
Cloned: Yes

10

**DNA Sequence:**

GGATCCATGATGTCTAAACTGGGAGTCTTGTGATCATCTGCTCTGCTTCTGTTCCCC  
TTACTGCTGTTCCGCTGGATGGAGATCAGCCTGCAGAGCGTACGCAGATCGAGCAG  
CATCCCTTGTGTTGACCAGAAAAGAAGGTGTTGCCGGTGGCCATGCCAGTAGATG  
15 CGGCATGGCTAGGTGTTGCTCGTCATGATAACGTGTTGATGANCGACCTCTCNAG  
(SEQ ID NO:270)

**Translation:**

MMSKLGVLLIICLLFPLTAVPLDGDQPAERTQIEQHPLFDQKRRCCRWPCCSRCGMAR  
20 CCFVMITC (SEQ ID NO:271)

**Toxin Sequence:**

Arg-Cys-Cys-Arg-Xaa4-Xaa3-Cys-Xaa3-Ser-Arg-Cys-Gly-Met-Ala-Arg-Cys-Cys-Phe-Val-Met-Ile-Thr-Cys-^ (SEQ ID NO:272)

25

Name: Pr3.1  
Species: parius  
Cloned: Yes

30

**DNA Sequence:**

GGATCCATGATGTCTAAACTGGGAGTCTTGTGACCATCTGCTCTGCTTCTGTTCCCC  
TTACTGCTCTCCGATGGATGGTATCAACCTGCAGACCGACTGTAGAGCGTATGC  
AGGACAAACATTCTGAGCAGCATCCCTCTTGA  
35 AAAAGAGAAGAGGAGGAGGCTGT  
TGCACACCTCCGAAGAAATGCAAAGACCGAGCCTGCAAACCTGCACGTTGCTGCGG  
CCCAGGATAACGTGTTGATGACCAACTTCTCGCC (SEQ ID NO:273)

**Translation:**

MMSKLGVLLTICLLFPLTALPMGDQPADRLVERMQDNISSEQHPFFEKRRGGCCTPP  
40 KKCKDRACKPARCCGPG (SEQ ID NO:274)

**Toxin Sequence:**

Arg-Gly-Gly-Cys-Cys-Thr-Xaa3-Xaa3-Lys-Lys-Cys-Lys-Asp-Arg-Ala-Cys-Lys-Xaa3-Ala-Arg-Cys-Cys-Gly-Xaa3-# (SEQ ID NO:275)

45

Name: Pr3.2  
Species: parius  
Cloned: Yes

5 **DNA Sequence:**

GGATCCATGATGTCTAAACTGGGAGTCTTGTGACCATCTGTCTGCTTCTGTTCCCC  
TTACTGCTCTCCGATGGATGGTATCAACCTGCAGACCGACTGTAGAGCGTATGC  
AGGACAACATTCTGAGCAGCATCCCTCTTGAAAAGAGAAGAGGGCTGTTGC  
ACACCTCCGAGGAAATGCAAAGACCGAGCCTGCAAACCTGCACGTTGTCGGCCC  
10 AGGATAACGTGTTGATGACCAACTTCTCGAG (SEQ ID NO:276)

## Translation:

MMSKLGVLLTICLLLFPALPMGDQPADRLVERMQDNISSEQHPFFEKRRGCCTPPR  
KCKDRACKPARCCGPG (SEQ ID NO:277)

15

**Toxin Sequence:**

Arg-Gly-Cys-Cys-Thr-Xaa3-Xaa3-Arg-Lys-Cys-Lys-Asp-Arg-Ala-Cys-Lys-Xaa3-Ala-Arg-  
Cys-Cys-Gly-Xaa3-# (SEQ ID NO:278)

20

Name: Ct3.1  
Species: coronatus  
Cloned: Yes

25

**DNA Sequence:**

GGATCCATGATGTCTAAACTGGGAGTCTTGTGACCATCTGTCTGCTTCTGTTCCAA  
TTACTGCCCTCCGCTGGATGAAGATCAACCTGCAGACCGACCTGCAGAGCGTATGC  
AGGACATTGCAACTGAACAGCATCCCTGTTGATCCGTCAAACGGTGCTGCGATT  
GGCCATGCATCCCAGGATGCACCCCTGTTGCTGCCTGATAACGTGTTGATGACC  
30 AACTTCTCGAG (SEQ ID NO:279)

## Translation:

MMSKLGVLLTICLLLFPITALPLDEDQPADRPAERMQDIATEQHPLFDPVKRCCDWPCIP  
GCTPCCLP (SEQ ID NO:280)

35

**Toxin Sequence:**

Cys-Cys-Asp-Xaa4-Xaa3-Cys-Ile-Xaa3-Gly-Cys-Thr-Xaa3-Cys-Cys-Leu-Xaa3-^ (SEQ ID  
NO:281)

40

Name: Ms3.1  
Species: musicus  
Cloned: Yes

45

**DNA Sequence:**

GGATCCATGATGTCTAAACTGGGAGTCTTGTGACCATCTGTCTGCTTCTGTTCCCTC  
TTCTGCTCTCCGATGGATGAAGATCAACCTGCAGACCTACCTGCAGAGCGTATGC

GGGACACTGCAACTGTAGATCATCCCTCCTATGATCCTGACAAAGCGTGCTGCGAG  
CAGAGCTGTACAACATGCTTCCGTGCTGCTAGCCTGAACACAGTAACGTGTTGAT  
GACCAACTTCTCGAG (SEQ ID NO:282)

5 **Translation:**

MMSKLGVLITICLLFPLSALPMDEDQLADLPAERMRTATVDHPSYDPDKACCEQSC  
TTCFPCC (SEQ ID NO:283)

**Toxin Sequence:**

10 Ala-Cys-Cys-Xaa1-Gln-Ser-Cys-Thr-Thr-Cys-Phe-Xaa3-Cys-Cys-^ (SEQ ID NO:284)

Name: bt3b  
Species: betulinus  
15 Isolated: Yes

**Toxin Sequence:**

Ala-Cys-Cys-Xaa1-Gln-Ser-Cys-Thr-Thr-Cys-Met-Xaa3-Cys-Cys-^ (SEQ ID NO:285)

20

Name: bt3c  
Species: betulinus  
Isolated: Yes

25 **Toxin Sequence:**

Cys-Cys-Xaa1-Gln-Ser-Cys-Thr-Thr-Cys-Met-Xaa3-Cys-Cys-Xaa4-? (SEQ ID NO:286)

30

Name: Pn3.2  
Species: pennaceus  
Cloned: Yes

**DNA Sequence:**

35 GGATCCATGATGTCTAAACTGGGAGTCTTGTGACCATCTGTCTGCTTCTGTTCCCC  
TTACTGCTCTTCCGCTGGATGGAGATCAACCTGCATACCAAGCTGCAGAGCGTATGC  
AGGCCGAGCATCATCCCTTGTGATCAGAAAAGACGGTGCTGCAAGTTCCATGCC  
CCGATAGTTGCAAATATTGTGTTGCGGGTGATGATAACATGTTGATGACCAACTT  
CTTGAG (SEQ ID NO:287)

40

**Translation:**

MMSKLGVLITICLLFPLTALPLGDQPAYQAAERMQAEHHPLFDQKRRCCFKPCPDS  
CKYLCG (SEQ ID NO:288)

45

**Toxin Sequence:**

Arg-Cys-Cys-Lys-Phe-Xaa3-Cys-Xaa3-Asp-Ser-Cys-Lys-Xaa5-Leu-Cys-Cys-# (SEQ ID NO:289)

5      Name:      Pu3.2  
Species:      pulicarius  
Cloned:      Yes

DNA Sequence:  
GGATCCATGATGTCTAAACTGGGAGTCTTGTGACCATCTGTCTGCTTGTGTTCCCC  
TTACTGCTCTTCCGATGGATGGTATCAACTTGCAGACCGACTGTAGAGCGTATGC  
10     AGGACAAACATTCATCTGAGCAGCATCCCTTCTTGATCCCGTCAAACGGTGTGCG  
TCAGCTGTTACATGGGATGCATCCCTTGTGCTTAGTAATAACGTGTTGATGACC  
AACTTCTCGAG (SEQ ID NO:290)

15     Translation:  
MMSKLGVLITCLLFPLTALPMGDQLADRLVERMQDNISSEQHPFFDPVKRCCVSCY  
MGCIPCCF (SEQ ID NO:291)

20     Toxin Sequence:  
Cys-Cys-Val-Ser-Cys-Xaa5-Met-Gly-Cys-Ile-Xaa3-Cys-Cys-Phe-^ (SEQ ID NO:292)

25     Name:      Pu3.3  
Species:      pulicarius  
Cloned:      Yes

DNA Sequence:  
GGATCCATGATGTCTAAACTGGGAGTCTTGTGACCGTCTGTCTGCTTGTGTTCCCC  
TTACTGCTCTTCCACTGGATGAAGATCAACTTGCAGACCGACCTGCAGAGCGTATGC  
30     AGGATGACACTTCAGCTGCACAGATTTCGGGTTGATCCCGTCAAACGGTGTGCA  
AATTGCTATGCTACTCGGGATGCACTCCTTGTGCCATATTGATAACGTGTTGATG  
ACCAACTTCTCGAG (SEQ ID NO:293)

35     Translation:  
MMSKLGVLTVCLLCPPLTALPLDEDQLADRPAERMQDDTSAAQIFGFDPVKRCCKLL  
CYSGCTPCCHI (SEQ ID NO:294)

Toxin Sequence:  
Cys-Cys-Lys-Leu-Leu-Cys-Xaa5-Ser-Gly-Cys-Thr-Xaa3-Cys-Cys-His-Ile-^ (SEQ ID NO:295)

40     Name:      Ra3.2  
Species:      rattus  
Cloned:      Yes

45     DNA Sequence:  
GGATCCATGATGTCTAAACTGGGAGTCTTGTGACCATCTGTCTGCTTGTGTTCCGC  
TTACTGCTCTTCCGATGGATGGTATCAACCTGCAGACCGACTGTAGAGCGTATAC

AGGACAAACATTCATCTGAGCAGCATCCCTCTTGAAAAGAGAAGAGGGCTGTGC  
GCACCTCCGAGGAAATGCAAAGACCGAGCCTGCAAACCTGCACGTTGCTGCGGCC  
AGGATAACGTGTTGATGACCAACTTCTCGAG (SEQ ID NO:296)

5 **Translation:**

MMSKLGVLLTICLLVFPLTALPMGDQPADRLVERIQDNISSEQHPFFEKRRGCCAPRK  
CKDRACKPARCCGPG (SEQ ID NO:297)

**Toxin Sequence:**

10 Arg-Gly-Cys-Cys-Ala-Xaa3-Xaa3-Arg-Lys-Cys-Lys-Asp-Arg-Ala-Cys-Lys-Xaa3-Ala-Arg-  
Cys-Cys-Gly-Xaa3-# (SEQ ID NO:298)

15 **Name:** Sm3.3

**Species:** stercusmuscarum

**Cloned:** Yes

**DNA Sequence:**

20 GGATCCATGATGTCTAAACTGGGAGTCTTGTGACAATCTGCTGCTCTGTTCCCC  
TTATTGCTCTCCGCTGGATGGAGATCAACCTGCAGACCGACCTGCAGAGCGTATGC  
AGGACGACATTTCATCTGAGAACGATCCCTGTTGATAAGAGACAACGGTGTGC  
AATGGGCGGAGGGGATGCTCCAGCAGATGGTGCAGAGATCACTCACGTTGTGCGG  
TCGACGATAACGTGTTGATGACCAACTTCTCGAG (SEQ ID NO:299)

25 **Translation:**

MMSKLGVLLTICLLFPLIALPLDGDQPADRPAERMQDDISSEKHPLFDKRQRCCNGRR  
GCSSRWCRDHSRCCGRR (SEQ ID NO:300)

**Toxin Sequence:**

30 Xaa2-Arg-Cys-Cys-Asn-Gly-Arg-Arg-Gly-Cys-Ser-Ser-Arg-Xaa4-Cys-Arg-Asp-His-Ser-Arg-  
Cys-Cys-# (SEQ ID NO:301)

35 **Name:** Eb3.1

**Species:** ebraeus

**Cloned:** Yes

**DNA Sequence:**

40 GGATCCATGATGTCTAAACTGGGAGTCTTGTGACCATCTGCTGCTCTGTTCCCC  
TTACTGCTCTCCACTGGATGAAGGTCAACCTGCAGACACTACCTGCAGAGCGTATGC  
AGGACATTGCAACTGAACAGCATCCCTGTTGATCCTGTCAAACGGTGTGCGAGC  
AGCCATGCTACATGGGATGCATCCCTGTTGCTTAATAAACGTGTTGATGACC  
AACTTCTCGAG (SEQ ID NO:302)

45 **Translation:**

MMSKLGVLLTICLLFPLTALPLDEGQPADLPAERMQDIATEQHPLFDPVKRCCEQPCY  
MGCIPCCF (SEQ ID NO:303)

**Toxin Sequence:**

Cys-Cys-Xaa1-Gln-Xaa3-Cys-Xaa5-Met-Gly-Cys-Ile-Xaa3-Cys-Cys-Phe-^ (SEQ ID NO:304)

5

**Name:** Eb3.2  
**Species:** ebraeus  
**Cloned:** Yes

10 **DNA Sequence:**

GGATCCATGATGTCTAAACTGGGAGTCTTGTGACCATCTGTCTGCTTCTGTTCCCC  
TTACTGCTCTTCCACTGGATGAAGATCAACCTGCAGACCTACCTGCAGAGCGTATGC  
AGGACATTGCAACTGAACAGCATCCCTGTTGATCCTGTCAAACGGTGTGCGCGC  
AGCCATGCTACATGGGATGCATCCCTGTTGCTTCTAATAATAACGTGTTGATGACC  
15 AACTTCTCGAG (SEQ ID NO:305)

**Translation:**MMSKLGVLLTICLLLFPPLTALPLDEDQPADLPAERMQDIATEQHPLFDPVKRCCAQPCY  
MGCIPCCF (SEQ ID NO:306)

20

**Toxin Sequence:**

Cys-Cys-Ala-Gln-Xaa3-Cys-Xaa5-Met-Gly-Cys-Ile-Xaa3-Cys-Cys-Phe-^ (SEQ ID NO:307)

25

**Name:** Fd3.2  
**Species:** flavidus  
**Cloned:** Yes

**DNA Sequence:**

30

GGATCCATGATGTCTAAACTGGGAGTCTTGTGACCATCTGTCTGCTTCTGTTCCCC  
TTACTGCTGTTCCGTTGGATGGAGATCAACCTGCAGACCCAGCCTGCAGAGCGTATGC  
AGAACGAGCAGCATCCCTGTTGATCAGAAAAGAAGGTGTGCTGCCGGTGGCCATGC  
CCCAGTATATGCGGCATGGCTAGGTGTTGCTCGTCATGATAACGTGTTGATGACCAA  
CTTTCTCGAG (SEQ ID NO:308)

35

**Translation:**MMSKLGVLLTICLLLFPPLTAVPLDGDQPADQPAERMQNEQHPLFDQKRRCCRWP CPSIC  
GMARCCSS (SEQ ID NO:309)

40

**Toxin Sequence:**Arg-Cys-Cys-Arg-Xaa4-Xaa3-Cys-Xaa3-Ser-Ile-Cys-Gly-Met-Ala-Arg-Cys-Cys-Ser-Ser-^  
(SEQ ID NO:310)

45

**Name:** Mf3.1  
**Species:** miliaris  
**Cloned:** Yes

**DNA Sequence:**

GGATCCATGATGTCTAAACTGGGAGTCTTGTGACCATCTGTCTGCTTCTGTTCCAA  
TTACTGCCCTTCCACTGGATGAAGATCAACCTGCAGACCGACCTGCAGAGCGTATGC  
5 AGGACATTGCAACTGAACAGCATTCCCTGTTGATCCGTCAAACGGTGTGCGATT  
GGCCATGCAGCGCAGGATGCTACCCTGCTTCCCTTAATAACGTGTGATGACC  
AACTNANGNAAAAAAA (SEQ ID NO:311)

**Translation:**

10 MMSKLGVLLTICLLLFPITALPLDEDQPADRPAAERMQDIATEQHPLFDPVKRCCDWPCS  
AGCYPCCFP (SEQ ID NO:312)

**Toxin Sequence:**

15 Cys-Cys-Asp-Xaa4-Xaa3-Cys-Ser-Ala-Gly-Cys-Xaa5-Xaa3-Cys-Cys-Phe-Xaa3-^ (SEQ ID  
NO:313)

Name: Mf3.2

Species: miliaris

20 Cloned: Yes

Notes:

**DNA Sequence:**

GGATCCATGATGTCTAAACTGGGAGTGGTGCCATTCTGCTTTCTGGTCCCTGTTCCCC  
25 TGGCAACACTCCAACACTGGATGCAGATCAACCTGCAGACCGACCTGCGCGTAAAAAG  
GGCATTGCAACTAAACGGCATCCCTGCTGATCCTGTCAGAGGGTGTGCCCTCCA  
ATGTGCACACCATGCTCCCTGCTGTTGTTAATAACGTGTGATGNATGATGN  
AN (SEQ ID NO:314)

30 Translation:

MMSKLGVPFVFLVLFPLATLQLDADQPADRPARKGIATKRHPLSDPVRGCCPPMCTPCFPCC  
FR (SEQ ID NO:315)

**Toxin Sequence:**

35 Gly-Cys-Cys-Xaa3-Xaa3-Met-Cys-Thr-Xaa3-Cys-Phe-Xaa3-Cys-Cys-Phe-Arg-^ (SEQ ID  
NO:316)

Name: Af3.1

40 Species: ammiralis

Cloned: Yes

**DNA Sequence:**

CAAGAGGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTGACCATCTG  
45 TCTGCTTCTGTTCCCTTAAGTGCCTTCCCGCTGGATGGAGATCAACCTGCAGACCA  
AGCTGCAGAGCGTATGCAGGCCGAGCAGCATCCCTGTTGATCAGAAAAGACGGT  
GTTGCAGGTTCCATGCCCGATACTTGCAGACATTGTGTTGCGGGTGATGATAAC

GTGCTGATGACCCACTTGTATCACGGCTACGTCAAGTGTCTAATGAATAAGTAAA  
ATGATTGCAGT (SEQ ID NO:317)

**Translation:**

5 MMSKLGVLLTICLLLFPALPLDGDQPADQAAERMQAEQHPLFDQKRRCRFPCTPDT  
CRHLCCG (SEQ ID NO:318)

**Toxin Sequence:**

10 Arg-Cys-Cys-Arg-Phe-Xaa3-Cys-Xaa3-Asp-Thr-Cys-Arg-His-Leu-Cys-Cys-# (SEQ ID  
NO:319)

Name: Af3.2

Species: ammiralis

15 Cloned: Yes

**DNA Sequence:**

20 CAAGAGGGATCGATAGCAGTTCATGATGTTAAACTGGGAGCTTGCTGACCATCTG  
TCTACTTCTGTTTCCCTTAATGCTGTTCCGCTGGATGGAGATCAACCTGCAGACCA  
ACCTGCAGAGCGTCTGCTGGACGACATTCTGAAAATAATCCCTTTATGATCC  
CGCCAAACGGTGTGCATGACTGCTCGTTGCACACCTTGTGGATGACCAGC  
CTCATCAAGTGTCTAACGAATAAGTAAAACGATTGCAGT (SEQ ID NO:320)

**Translation:**

25 MMFKLGVLLTICLLLFSLNNAVPLDGDQPADQPAERLDDISSENNPFYDPAKRCCMTCF  
GCTPCCG (SEQ ID NO:321)

**Toxin Sequence:**

30 Cys-Cys-Met-Thr-Cys-Phe-Gly-Cys-Thr-Xaa3-Cys-Cys-# (SEQ ID NO:322)

Name: Af3.3

Species: ammiralis

35 Cloned: Yes

**DNA Sequence:**

40 CAAGAAGGATCGATAGCAGTTCATGATGTTAAACTGGGAGCCCTGTTGACCATCT  
GTCTACTTCTGTTTCCCTTAATGCTGTTCCGCTGGATGGAGATCAACATGCAGACC  
AACCTGCAGAGCGTCTGCTGGACGCCCTCAACTGAAAATCATCCCTATATGATC  
CCGTCAAACGGTGTGCATGATTGCGACTATTCTGCTGGCCTGCTGTA  
TTTTTCATAACCTTGTATCGCGGCCTCATCCTAGTGTCAAATGAATAAGTAAAA  
CGATTGCAGT (SEQ ID NO:323)

**Translation:**

45 MMSKLGALLTICLLLFSLTAVPLDGDQHADQPAERLQDRLPTENHPLYDPVKRCCDDSE  
CDYSCWPCCIFS (SEQ ID NO:324)

**Toxin Sequence:**

Cys-Cys-Asp-Asp-Ser-Xaa1-Cys-Asp-Xaa5-Ser-Cys-Xaa4-Xaa3-Cys-Cys-Ile-Phe-Ser-^ (SEQ ID NO:325)

5

Name: Af3.4  
Species: ammiralis  
Cloned: Yes

10

**DNA Sequence:**

CAAGAGGGATCGATAGCAGTTCATGATGTTAAACTCGGAGTCCTGCTGACCATCTG  
TCTACTTCTGTTTCCCTAATGCTGTTCCGCTGGATGGAGATCAACATGCAGACCAA  
CCTGCAGAGCGTCTGCAGGACCCTCAACTGAAAATCATCCCTTATATGATCCC  
15 GTCAAACCGGTGTTGCAGGTTGTTATGCCTCAGTTGCAACCCCTGTTGTGGATGACCA  
GCTTGTATCACGGCCTCATCAAGTGTCTAATGAATAAGTAAAACGATTGCAGT  
(SEQ ID NO:326)

**Translation:**

20

MMFKLGVLITICLLFSLIAVPLDGDQHADQPAERLQDRLPTEHPLYDPVKRCCRLLC  
LSCNPCCG (SEQ ID NO:327)

**Toxin Sequence:**

Cys-Cys-Arg-Leu-Leu-Cys-Leu-Ser-Cys-Asn-Xaa3-Cys-Cys-# (SEQ ID NO:328)

25

Name: Af3.6  
Species: ammiralis  
Cloned: Yes

30

**DNA Sequence:**

CAAGAAGGATCGATAGCAGTTCATGATGTCATAACTGGAGCCTGTTGACCATCT  
GTCTACTTCTGTTTCCCTTACTGCTGTTCCGCTGGATGGAGATCAACATGCAGACC  
AACCTGCAGAGCGTCTGCAGGACCCTCAACTGAAAGATCATCCCTTATTGATC  
35 CCAACAAACGGTGTGCGATGATTGGAATGCGGCTATTGATGCTGGCCTGCTGTT  
ATGGATAAGCTTGTATCGCGGCCATCCAGTGTCAACGAATAAGTAAAACGATT  
GCAGT (SEQ ID NO:329)

**Translation:**

40

MMSKLGALLTICLLFSLIAVPLDGDQHADQPAERLQDRIPTEHPLFDPNKRCCDDSE  
CGYSCWPCCY (SEQ ID NO:330)

**Toxin Sequence:**

Cys-Cys-Asp-Asp-Ser-Xaa1-Cys-Gly-Xaa5-Ser-Cys-Xaa4-Xaa3-Cys-Cys-Xaa5-# (SEQ ID NO:331)

45

Name: Sf3.1  
Species: spurius  
Cloned: Yes

5 **DNA Sequence:**  
CAAGAAGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGCTGACCATCT  
GTCTGCTTCTGTTCCACGTACTTCTCTTCCGCTGGATGGAGATCAACCTGCAGTCCG  
ATCTGCAAAGCGTATGCATTCATCTATACAGCGTCGTTCTTGATCCCGTCAAACG  
10 GTGTTGCCCTAGATGCAGCGAGTGCAACCCCTGTTGGATGACCAGCTTGTCAATC  
GCGGCCTCATTAAGTGTCTAATGAATAAGTAAAATGATTGCAGT (SEQ ID NO:332)

**Translation:**

MMSKLGVLLTICLLLFPRTSLPLGDQPAVRSAKRMHSSIQRFFDPVKRCCPRCSECNP  
CCG (SEQ ID NO:333)

15 **Toxin Sequence:**  
Cys-Cys-Xaa3-Arg-Cys-Ser-Xaa1-Cys-Asn-Xaa3-Cys-Cys-# (SEQ ID NO:334)

20 Name: Om3.1  
Species: omaria  
Cloned: Yes

**DNA Sequence:**

25 CAAGAGGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTCGTTGACCATCT  
GTCTACTTCTATTTCCCTTACTGCTGTTCCGCTTGATGGAGATCAACATGCAGACCCA  
ACCTGCAGAGCGTCTGCAGGGCGACATTATCTGAAAAGCATCCCTATTTAATCC  
CGTCAAACGGTGTGCGATGAGGAAGAATGCAGCAGTGCTGGCCTTGTGTT  
30 GGGGGTGATCAGCTTGTATCGCGGCCTCATCAAGTGTCTAATGAATAAGTAAAAT  
GATTGCAGT (SEQ ID NO:335)

**Translation:**

MMSKLGVSLLTICLLLFSLTAVPLGDQHADQPAERLQGDILSEKHPLFNPVKRCCDEEE  
CSSACWPCCWG (SEQ ID NO:336)

35 **Toxin Sequence:**  
Cys-Cys-Asp-Xaa1-Xaa1-Xaa1-Cys-Ser-Ser-Ala-Cys-Xaa4-Xaa3-Cys-Cys-Xaa4-# (SEQ ID NO:337)

40 Name: Om3.2  
Species: omaria  
Cloned: Yes

45 **DNA Sequence:**  
CAAGAAGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTTGATCATCTG  
TCTACTTCTGTCCTTACTGCTGTTGGAGGATGGAGATCAACCTGCAGACCG

ACCTGCAGAGCGTATGCAGGACGACATTCAACTGAGCATCATCCCTTTATGATCC  
 CGTCAAACGGTGTGCAAGTACGGGTGGACATGCTGCTAGGATGCACTCCTGTGA  
 TTGTTGACCAGTTGTATCGCGGCCTCGTCAAGTGTCTAATGAATAAGTAAAACG  
 ATTGCAGT (SEQ ID NO:338)

5

**Translation:**

MMSKLGVLIIICLLCPLTAVLEDGDQPADRPAERMQDDISTEHHPFYDPVKRCKY  
 WTCLLGCTPCDC (SEQ ID NO:339)

10

**Toxin Sequence:**

Cys-Cys-Lys-Xaa5-Gly-Xaa4-Thr-Cys-Leu-Leu-Gly-Cys-Thr-Xaa3-Cys-Asp-Cys-^ (SEQ ID NO:340)

15

**Name:** Om3.3  
**Species:** omaria  
**Cloned:** Yes

**DNA Sequence:**

20

CAAGAGGGATCGATAGCAGTTCATGATGTCTATACTGGGAGTCTTGTGATCATCTG  
 TCTACTTCTGTGTCCTTACTGCTGTTCTGGAGGATGGAGATCAACCTGCAGACCG  
 ACCTGCAGAGCGTATGCAGGACGGCATTTCATCTGAACATCATCCCTTTGGATCC  
 CGTCAAACGGTGTGCCATCTATGGCATGCCCTTGGATGCTCGCCTTGTGTTG  
 GTGACCAGCTTGTATCGCGGCCTCATCAAGTGTCTAATGAATAAGTAAAACGATT  
 25 GCAGT (SEQ ID NO:341)

**Translation:**

MMSILGVLLIICLLCPLTAVLEDGDQPADRPAERMQDGISSEHHPFLDPVKRCCHLLAC  
 RFGCSPCCW (SEQ ID NO:342)

30

**Toxin Sequence:**

Cys-Cys-His-Leu-Leu-Ala-Cys-Arg-Phe-Gly-Cys-Ser-Xaa3-Cys-Cys-Xaa4-^ (SEQ ID NO:343)

35

**Name:** Om3.4  
**Species:** omaria  
**Cloned:** Yes

**DNA Sequence:**

40

CAAGAAGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTGATCATCTG  
 TCTACTTCTTGTCCTTACTGCTGTTCCGCAGGATGGAGATCAACCTGCAGACCG  
 ACCTGCAGAGCGTATGCAGGGCGGCATTTCATCTGAACATCATCCCTTTGGATCC  
 CGTCAAACGGTGTGCAGGTACGGGTGGACATGCTGGCTAGGATGCACTCCCTGTG  
 GTTGTGACCAGCTTGTATCGCGGCCTCATCAAGTGTCTAATGAATAAGTAAAAC  
 45 GATTGCAGT (SEQ ID NO:344)

**Translation:**

MMSKLGVLIIICLLCPLTAVPQDGDQPADRPAERMQGGISSEHPFFDPVKRCCRYGW  
TCWLGCTPCGC (SEQ ID NO:345)

**Toxin Sequence:**

5 Cys-Cys-Arg-Xaa5-Gly-Xaa4-Thr-Cys-Xaa4-Leu-Gly-Cys-Thr-Xaa3-Cys-Gly-Cys-^ (SEQ ID NO:346)

10 Name: Ep3.1  
Species: episcopatus  
Cloned: Yes

**DNA Sequence:**

15 CAAGAAGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTGACCATCTG  
TCTACTTCTGTTTCCCTTATTGCTGTTCCGCTTGATGGAGATCAACATGCAGACCAA  
CCTGCAGAGCGTCTGCAGGGCGACATTTATCTGAAAAGCATCCCTATTTATGCCT  
GTCAAACGGTGTGCGATGAGGACGAATGCAACAGTTCATGCTGGCCTTGTGTTGG  
GGGTGATCAGCTTGTATCGCGGCCTGATCAAGTGTATAATGAATAAGTAAAACG  
ATTGCAGT (SEQ ID NO:347)

20 **Translation:**  
MMSKLGVLLTICLLFSLIAVPLGDQHADQPAERLQGDILSEKHPLFMPVKRCCDEDE  
CNSSCWPCCWG (SEQ ID NO:348)

25 **Toxin Sequence:**  
Cys-Cys-Asp-Xaa1-Asp-Xaa1-Cys-Asn-Ser-Ser-Cys-Xaa4-Xaa3-Cys-Cys-Xaa4-# (SEQ ID NO:349)

30 Name: Ep3.2  
Species: episcopatus  
Cloned: Yes

**DNA Sequence:**

35 CAAGAGGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTGACCATCTG  
TCTACTTCTGTTTCCCTTATTGCTGTTCCGCTTGATGGAGATCAACATGCAGACCAA  
CCTGCAGAGCGTCTGCAGGGCGACATTTATCTGAAAAGCATCCCTATTTATGCCT  
GTCAAACGGTGTGCGATGAGGACGAATGCAAGCAGTTCATGCTGGCCTTGTGTTGG  
GGATGAGCAGCTTGTATCGCGGCCTCATCAAGTGTCTAATGAATAAGTAAAACG  
40 ATTGCAGT (SEQ ID NO:350)

**Translation:**  
MMSKLGVLLTICLLFSLIAVPLGDQHADQPAERLQGDILSEKHPLFMPVKRCCDEDE  
CSSSCWPCCWG (SEQ ID NO:351)

45 **Toxin Sequence:**

Cys-Cys-Asp-Xaa1-Asp-Xaa1-Cys-Ser-Ser-Ser-Cys-Xaa4-Xaa3-Cys-Cys-Xaa4-# (SEQ ID NO:352)

5   **Name:**    Ep3.3  
  **Species:**   episcopatus  
  **Cloned:**    Yes

**DNA Sequence:**

10   CAAGAGGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTGACCATCTG  
      TCTACTTCTGTTTCCCTTACTGCTGTTCCGCTTGATGGAGATCAACATGCAGACCAA  
      CCTGCAGAGCGTCTGCAGGGCAGACATTATCTGAAAAGCATCCCTATTTAATCCC  
      GTCAAACGGTGTGCCCGCGGCCATGTGCCATGGGATGCAAGCCTTGTGTTGG  
      ATGAGCAGCTTGTTATCGTGGCCTCATCAAGTGTCTAATGAATAAGTAAAACGATT  
15    GCAGT (SEQ ID NO:353)

**Translation:**

MMSKLGVLLTICLLLFSLTAVPLGDQHADQPAERLQGDILSEKPLFNPVKRCCPAAA  
CAMGCKPCCG (SEQ ID NO:354)

20   **Toxin Sequence:**  
      Cys-Cys-Xaa3-Ala-Ala-Ala-Cys-Ala-Met-Gly-Cys-Lys-Xaa3-Cys-Cys-# (SEQ ID NO:355)

25   **Name:**    Au3.2  
  **Species:**   aulicus  
  **Cloned:**    Yes

**DNA Sequence:**

30   CAAGAGGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTGACCATCTG  
      TCTGCTTCTGTTCCGTTACTGCTCTTCCGCCGGATGGAGATCAACCTGCAGACCG  
      AGCTGCAGAGCGTAGGCAGGTGAGCAGCAGATCCCGTGTGATCATGAAAGAGGGT  
      GTTGCTGCCACCATGCCACAGTATTGCGCTGCTTCTGTTGCCGGTGTGATAAC  
      GTGTTGATGACCCACTTGTATCACGGCTGCGTCAAGTGTCTAATGAATAAGTAAA  
35    ATGATTGCACT (SEQ ID NO:356)

**Translation:**

MMSKLGVLLTICLLLFSVTALPPDGDQPADRAAERRQVEQHPVFDHERGCCSPPCHSIC  
AAFCCG (SEQ ID NO:357)

40   **Toxin Sequence:**  
      Gly-Cys-Cys-Ser-Xaa3-Xaa3-Cys-His-Ser-Ile-Cys-Ala-Ala-Phe-Cys-Cys-# (SEQ ID NO:358)

45   **Name:**    Au3.3  
  **Species:**   aulicus  
  **Cloned:**    Yes

**DNA Sequence:**

CAAGAGGGATCGATAGCAGTTCATGATGTCATAACTGGGAGTCTTGTGACCATCTG  
TCTACTTCTGTTTCCCTTACTGCTGTTCCGCTTGATGGAGATCAACATGCAGACCAA  
5 CCTGCAGAGCGTCTGCAGGGCAGACATTATCTGAAAAGCATCCCTATTAATCCC  
GTCAAACGGTGTGCCGACCCTGGCATGCCATGGATGCAAGCCTTGTGTTGG  
ATGAGCAGCTTGTATCGTGGCCTCATCAAGTGTCTAATGAATAAGTAAAATGATT  
GCAGT (SEQ ID NO:359)

**10 Translation:**

MMSKLGVLITICLLFSLTAVPLDGDQHADQPAERLQGDILSEKHPLFNPVKRCCRPVA  
CAMGCKPCCG (SEQ ID NO:360)

**Toxin Sequence:**

15 Cys-Cys-Arg-Xaa3-Val-Ala-Cys-Ala-Met-Gly-Cys-Lys-Xaa3-Cys-Cys-# (SEQ ID NO:361)

**Name:** Au3.4

**Species:** aulicus

20 **Cloned:** Yes

**DNA Sequence:**

CAAGAGGGATCGATAGCAGTTCaTGATGTCATAACTGGGAGTCTTGTGATCATCTG  
TCTACTTCTGCTCCCTTACTGCTGTTCCGCTGGATGGAGATCAACCTGCAGACCG  
25 ACCTGCAGAGCGTATGCAGGACGACATTTCATCTGAACATCAACCCATGTTGATGC  
CATCAGACAGTGTGCCCCGGCGGTGGCATGCCATGGATGCGAGCCTTGTGTTG  
GATGACCAGCTTGTATCGCGGCCATCAAGTGTCTAATGAATAAGTAAAATGAT  
TGCAGT (SEQ ID NO:362)

**30 Translation:**

MMSKLGVLIIICLLSPLTAVPLDGDQPADRPAERMQDDISSEHQPMFDAIRQCCPAVA  
CAMGCEPCCG (SEQ ID NO:363)

**Toxin Sequence:**

35 Xaa2-Cys-Cys-Xaa3-Ala-Val-Ala-Cys-Ala-Met-Gly-Cys-Xaa1-Xaa3-Cys-Cys-# (SEQ ID NO:364)

**Name:** Ae3.1

40 **Species:** aureus

**Cloned:** Yes

**DNA Sequence:**

CAAGAAGGGATCGATAGCAGTTCATGATGTCATAACTGGGAGCCTTGTGACCATCT  
45 GTCTACTTCTGTTTCCCTTACTGCTGTTCCGCTGGATGGAGATCAACATGCAGACCC  
AACATGCAGAGCGTCTGCATGACCGCCTCCAATGAAAATCATCCCTATATGATC  
CCGTCAAACGGTGTGCGATGATCGGAATGCGACTATTCTGCTGGCCTTGCTGTA

TTTTGGATAACCTTGTATCGCGGCCTCATCAAGTGTCAAATGAATAAGTAAAAC  
GATTGCAGT (SEQ ID NO:365)

**Translation:**

5 MMSKLGALLTICLLLFSLTAVPLGDQHADQHAERLHDRLPTENHPLYDPVKRCCDDS  
ECDYSCWPCCIFG (SEQ ID NO:366)

**Toxin Sequence:**

10 Cys-Cys-Asp-Asp-Ser-Xaa1-Cys-Asp-Xaa5-Ser-Cys-Xaa4-Xaa3-Cys-Cys-Ile-Phe-# (SEQ ID  
NO:367)

Name: Ae3.2

Species: aureus

15 Cloned: Yes

**DNA Sequence:**

20 CAAGAGGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGCCTTGTGACCATCT  
GTCTACTTCTGTTCCCTAACTGCTGTCGCTGGATGGAGATCAACATGCAGACC  
AACCTGCAGAGCGTCTGCAGGACCGCATTCCAAGTAAAATCATCCCTTATTGATC  
CGAACAAACGGTGTGCAATGATTGGAAATGCGACGATTGCTGGCCCTGCTGTT  
ATGGATAACCTTGTATCGCGGCCTCATCAAGTGTCAAATGAATAAGTAAAACGAT  
TGCAGT (SEQ ID NO:368)

25 **Translation:**

MMSKLGALLTICLLLFSLTAVPLGDQHADQPAERLQDRIPTENHPLFDPNKRCCNDWE  
CDDSCWPCCYHG (SEQ ID NO:369)

**Toxin Sequence:**

30 Cys-Cys-Asn-Asp-Xaa4-Xaa1-Cys-Asp-Asp-Ser-Cys-Xaa4-Xaa3-Cys-Cys-Xaa5-# (SEQ ID  
NO:370)

Name: Cn3.1

Species: consors

35 Cloned: Yes

**DNA Sequence:**

40 CAAGAGGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTGACCATCTG  
TTGCTTCTGTTCCCTTAAGTGCCTTCAATGGATGGAGATCAATCTGTAGACCGA  
CCTGCAGAGCGTATGCAGGACACATTCTAGCTGAGCTGCATCCCTGTCATCAG  
AAAAGAATGTGTTGCGCGAAGGTGCGCCATGCCCCAGCTATTCAGAAACAGTCA  
GATTGTCATTGTTAAATGACAACGTGTCGATGACCAACTCGTTATCAGCACT  
AATGAATAAGTAAAATGATTGCAGT (SEQ ID NO:371)

45

**Translation:**

MMSKLGVLITCLLFPLTALPMGDQSVDRPAERMQDDISSELHPLFNQKRMCCGEG  
APCPSYFRNSQICHCC (SEQ ID NO:372)

**Toxin Sequence:**

5 Met-Cys-Cys-Gly-Xaa1-Gly-Ala-Xaa3-Cys-Xaa3-Ser-Xaa5-Phe-Arg-Asn-Ser-Gln-Ile-Cys-His-  
Cys-Cys-^ (SEQ ID NO:373)

10 **Name:** Cn3.3  
**Species:** consors  
**Cloned:** Yes

**DNA Sequence:**

15 TAAGAGGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTGACCATCTG  
TCTGCTTCTGTTCCCTTATTGCTCTTCCAATGGATGGAGATCAACCTGCAGACCGA  
CCTGCAGAGCGTATGCAgGACGACATTCTCAGCAGCATCCCTTGTGATAAG  
AGAGGCCGCTGTTGCGATGTGCCGAACGCATGCTCCGGCAGATGGTGCAGAGATCA  
CGCACAAATGTTGCGGATGACGATAACGTGTGATGACCAACTTGTGATCACGGCTA  
CATCAAGTGAATAAGTAAAACGATTGCAGT (SEQ ID NO:374)

20 **Translation:**

MMSKLGVLITCLLFPLIALPMGDQPADRPAERMQDDISSEQHPLFDKRGRCVDVPN  
ACSGRWCRDHAQCCG (SEQ ID NO:375)

25 **Toxin Sequence:**

Gly-Arg-Cys-Cys-Asp-Val-Xaa3-Asn-Ala-Cys-Ser-Gly-Arg-Xaa4-Cys-Arg-Asp-His-Ala-Gln-  
Cys-Cys-# (SEQ ID NO:376)

30 **Name:** Cn3.4  
**Species:** consors  
**Cloned:** Yes

**DNA Sequence:**

35 CAAGAGGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTGACTGTCTG  
TTTGCTTCTGTTCCCTTACTGCTCTTCCGATGGATGGAGATCAACCTGCAGACCAA  
CCTGCAGAGCGTATGCAAGGACGACATTCTCAGCAGCATCCCTTGTGATAAG  
AGACAAAGGTGTTGCACTGGGAAGAAGGGGTATGCTCCGGTAAAGCATGCAAAA  
GTCTCAAATGTTGCTCTGGACGATAACGTGTGATGACCAACTTGTATCACGGCT  
40 ACGTCAAGTGTCTAGTGAATAAGTAAAACGATTGCAGT (SEQ ID NO:377)

**Translation:**

MMSKLGVLITVCLLFPLTALPMGDQPADQPAERMQDDISSEQHPLFDKRGRCCTGK  
KGSCSGKACKSLKCCSGR (SEQ ID NO:378)

45 **Toxin Sequence:**

Xaa2-Arg-Cys-Cys-Thr-Gly-Lys-Lys-Gly-Ser-Cys-Ser-Gly-Lys-Ala-Cys-Lys-Ser-Leu-Lys-Cys-Cys-Ser-# (SEQ ID NO:379)

5      Name:      Em3.1  
 Species:      emaciatus  
 Cloned:      Yes

**DNA Sequence:**

10      CAAGAGGGATCGATAGCAGTTCATGATGTCTAAACTGGGAGTCTTGTGACCATCTGTCCTGCTTGTGTT  
 TCCCCCTTACTGTTCTCCGATGGATGGAGATCAACCTGCAGACCTACCTGCATTGCGTGCAGTCTT  
 TGCACCTGAACATAGTCCCCGGTTGACCCCGTCAAACGGTGTGCTGCGGGATTGCAGTGTTGCAT  
 CCCCTGTTGCCCGTATGGATCACCTGATTATTGCGGCCACGTCAAGTGTCTAATGAATAAGTAAAATG  
 ATTGCAGT (SEQ ID NO:380)

15      **Translation:**

MMSKLGVLLTICLLLFPPLTVLPMDQPADLPALRAQFFAPEHSPRFDPVKRCCSRDCSVCIPCCPYGSP  
 (SEQ ID NO:381)

20      **Toxin Sequence:**

Cys-Cys-Ser-Arg-Asp-Cys-Ser-Val-Cys-Ile-Xaa3-Cys-Cys-Xaa3-Xaa5-Gly-Ser-Xaa3-^ (SEQ  
 ID NO:382)

25      **Where:**

Xaa1 is Glu or  $\gamma$ -carboxy-Glu

Xaa2 is Gln or pyro-Glu

Xaa3 is Pro or hydroxy-Pro

30      Xaa4 is Trp or bromo-Trp

Xaa5 is Tyr,  $^{125}$ I-Tyr, mono-iodo-Tyr, di-iodo-Tyr, O-sulpho-Tyr or O-phospho-Tyr

^ is free carboxyl or amidated C-terminus, preferably free carboxyl

# is free carboxyl or amidated C-terminus, preferably amidated

? = Status of C-term not known.

35

TABLE2

Alignment of  $\mu$ -Conopeptides (SEQ ID NO:)

TYPE 1

40	A3.4 (F283)	---CCKVQ-CES--C---TPCC^ (383)
	Ak3.1 (F585)	---CCELP-CGPGFC---VPCC^ (384)
	Ar3.1	---CCERP-CNIG-C---VPCC^ (385)
	Bn3.1 (F586)	---CCNWP-CSMG-C---IPCCYY^ (386)
	Bt3.1	---CCELP-CH-G-C---VPCCWP^ (387)
45	Bt3.2	---CCGLP-CN-G-C---VPCCWPS^ (388)
	Bt3.3	---CCSRN-CAV--C---IPCCPNWPA^ (389)
	bt3a	---CCKQS-CTT--C---MPCCW^ (390)

	bt3b	--ACCXQS-CTT--C---MPCC^ (391)
	bt3c	--CCEQS-CTT--C---MPCCW? (392)
	Ca3.3	R--CCRYP-CPDS-C--HGSCCYK^ (393)
	Ca3.4	--CCPPVACNMG-C---KPCC# (394)
5	Ca3.5	--CCDDSECDYS-C---WPCCMF# (395)
	Ca3.6 (F349)	--CCRR--CYMG-C---IPCCF^ (396)
	Circling	--CCPPVACNMG-C---KPCCG^ (397)
	Comatose/Death	SKQCCHLAAACRFG-C---TOCCN^ (398)
10	Cp3.1 (F594)	S---CCR--DCGED-C---VGCCR^ (399)
	Ct3.1 (Z726)	--CCDWp-CIPG-C---TPCCLP^ (400)
	Da3.1	--CCDDSECDYS-C---WPCCILS^ (401)
	Da3.2	Z-QCCPPVACNMG-C---EPCC# (402)
	Da3.3	--CCNAGFCRFG-C---TPCCW^ (403)
	Di3.1	Z---CCVHP-C-P--C---TPCCR^ (404)
15	Fi3.1	--CCPWP-CNIG-C---VPCC^ (405)
	Fi3.2	--CCSKN-CAV--C---IPCCP^ (406)
	Fi3.3	--CCRWP-CP-ARC---GSCCL^ (407)
	Fi3.4	--CCELSRCL-G-C---VPCCTS^ (408)
	Fi3.5	--CCELSKCH-G-C---VPCCIP^ (409)
20	Ge3.1 (F590)	Z---CCTF--CNFG-C---QPCCVP^ (410)
	Ge3.2 (F343/Z734)	Z---CCTF--CNFG-C---QPCCLT^ (411)
	Ge3.3 (F590)	Z---CCTF--CNFG-C---QPCCVP^ (412)
	Gm3.1	--CCDDSECDYS-C---WPCCMF# (413)
	Gm3.2	G---CCHLLAACRFG-C---SPCCW^ (414)
25	Gm3.3	--CCSWDVCDHPSC---T-CCG# (415)
	La3.1	--CCDWp-CS-G-C---IPCC^ (416)
	Lp3.1 (F340)	ZINCCPWP-CPST-C---RHQCCH^ (417)
	Lv3.1 (F341)	ZINCCPWP-CPDS-C---HYQCCH^ (418)
	Mr3.2	--CCRRLS-CGLG-C---HPCC# (419)
30	Mr3.3	--ECCGSFACRFG-C---VPCCV^ (420)
	Mr3.4	SKQCCHLPACRFG-C---TPCCW^ (421)
	Mr3.5 (F286)	-MGCCPFP-CKTS-C---TTLCC# (422)
	Ms3.1 (Z738)	--ACCEQS-CTT--C---FPCC^ (423)
	Nb3.1 (F87)	--CCELP-CGPGFC---VPCC^ (424)
35	Pu3.1 (F339)	--CCN-S-CYMG-C---IPCCF^ (425)
	Qc3.1 (F342)	ZR-CCQWP-CPGS-C---RCCRT# (426)
	Qc3.2	ZR-CCRWP-CPGS-C---RCCRYR^ (427)
	Qc3.3	R---CCRYP-CPDS-C---HGSCCYK^ (428)
	QcIIIA	--CCSQD-CLV--C---IOCCPN# (429)
40	QcIIIB	--CCSRH-CWV--C---IOCCPN? (430)
	Ra3.1 (F351)	Z-TCCS-N-CGED-C---DGCCQ^ (431)
	Scratcher I	--CCR-T-C-FG-C---TOCC# (433)
	Ts3.1 (F592)	--CCH-K-CYMG-C---IPCCI^ (434)
	Ts3.2 (F345)	K---CCRPP-CAMS-C-GMARCCY^ (435)
45	Bt3.5 (Z495)	R---CCRWP-CPSI-C-GMARCCFVMITC^ (436)
	Bt3.6 (Z497)	R---CCRWP-CP-SRC-GMARCCFVMITC^ (437)
	Tx3.1	F---CCDSNWCHISDC---ECCY# (438)

U014	---CCHWNWCDHL-C---SCCGS^ (439)
U017	--DCCOLPACPFG-C---NOCC# (440)
U019	--CCAPSACRLG-C---ROCCR^ (441)
U020	--CCAOSACRLG-C---ROCCR^ (442)
5 U022	--CCAPSACRLG-C---RPCCR^ (443)
U024	--GCCGSFACRFG-C---VOCCV^ (444)
U031	--CCSWDVCDHPSC---TCC# (445)
U032 (F353)	R--CCKFP-CPDS-C--RYLCC# (446)
Ae3.1	--CCDDSECDYS-C---WPCCIF# (447)
10 Ae3.2	--CCNDWECDDS-C---WPCCY# (448)
Af3.1	R--CCR-FPCPDT-C---RHLCC# (449)
Af3.2	--CC---MTC-FG-C---TPCC# (450)
Af3.3	--CCDDSECDYS-C---WPCCIFS^ (451)
Af3.4	--CCR-LLC-LS-C---NPCC# (452)
15 Af3.6	--CCDDSECGYS-C---WPCCY# (453)
Au3.2	G--CCS-PPCHSI-C---AAFCC# (454)
Au3.3	--CCRPVACAMG-C---KPCC# (455)
Au3.4	Z--CCPAVACAMG-C---EPCC# (456)
Em3.1	--CCS-RDC-SV-C---IPCCPYGSP^ (457)
20 Ep3.1	--CCDEDECNSS-C---WPCCW# (458)
Ep3.2	--CCDEDECSSS-C---WPCCW# (459)
Ep3.3	--CCPAACAMG-C---KPCC# (460)
Om3.1	--CCDEECSSA-C---WPCCW# (461)
Om3.3	--CCHLLACRFG-C---SPCCW^ (462)
25 Sf3.1	--CC---PRC-SE-C---NPCC# (463)

TYPE 2

30 Pn3.2 (AA049)	-RCC--KFP-CPDS-C--KYLCC# (464)
Fd3.2 (Z831)	-RCC--RWP-CPSI-C-GMARCCSS^ (465)
Pu3.3 (AA405)	--CC--KLL-CYSG-C---TPCCHI^ (466)
Eb3.1 (Z821)	--CC--EQP-CYMG-C---IPCCF^ (467)
Eb3.2 (Z822)	--CC--AQP-CYMG-C---IPCCF^ (468)
35 Pu3.2 (AA403)	--CC--V-S-CYMG-C---IPCCF^ (469)
Mf3.1 (Z882)	--CC--DWP-CSAG-C---YPCCFP^ (470)
Mf3.2 (Z885)	-GCC--PPM-C-TP-C---FPCCFR^ (471)
Ra3.2 (AA414)	RGCCAPPRK-CKDRACK-PARCCGP# (472)
Sm3.3 (AA419)	ZRCCNGRRG-CSSRWC RDHSRCC# (473)
40 Cn3.3	GRCCDVPNA-CSGRWCRDHAQCC# (474)
Cn3.4	ZRCCTGKGSCSGKACKSL-KCCS# (475)

TYPE 3

45 A3.1	-MCCGEGRKCP SYFRNSQICCHCC^ (476)
A3.2 (F84)	--CCR--WPCPRQIDGEY-CGCC# (477)
Bu3.5	-RCCGEGLTCPRYWKNSQICACC^ (478)
Ca3.1	--CCGGPGSCP VYFRDNFICGCC^ (479)

Cr3.1	RKCCGKDGPCPKYFKDNF1CGCC^ (480)
E3.1	--CCS--WPCPRYSNGKLVCFCCL# (481)
M3.2	--CCGPGGSCPVYFRDNF1CGCC^ (482)
M3.3	-MCCGESAPCPSPYFRNSQ1CHCC^ (483)
5 M3.4	ZKCCGPGGSCPVYFTDNF1CGCC^ (484)
M3.5	ZKCCGPGGSCPVYFRDNF1CGCC^ (485)
S3.1	ZKCCGEGSSCPKYFKNNF1CGCC^ (486)
U001	ZKCCS-GGSCPPLYFRDRLICPCC^ (487)
U034	ZKCCGPGASCPRYFKDNF1CGCC^ (488)
10 Cn3.1	-MCCGEGAPCPSPYFRNSQ1CHCC^ (489)

TYPE 4

15 A3.3 (F83)	ZK--CCTGK---KGCSGKACKNL-KCCS# (490)
A3.5 (Z488)	ZK--CCTGR---KGCSGKACKNL-KCCS# (491)
Bu3.1	VTDRCCK---GKREC-GRWCRDHSRCC# (492)
Bu3.1A	VGDRCCK---GKRGC-GRWCRDHSRCC# (493)
Bu3.2	VGERCCK---NGKRGC-GRWCRDHSRCC# (494)
20 Bu3.3	IVDRCCN-KGNGKRGC-SRWCRDHSRCC# (495)
Bu3.4	VGLYCCRPKPNGQMMC-DRWCEKNSRCC# (496)
Ca3.2	-RD-CCTPP---KK-CKDRQCKPQ-RCCA# (497)
L3.1	GRD-CCTPP---RK-CKDRACKPQ-RCCG# (498)
L3.2	ZRL-CCGFP---KS-CKRSRQCKPH-RCC# (499)
25 La3.2	-RD-CCTPP---KK-CKDRQCKPA-RCCG# (500)
La3.3	RPP-CCTYD---GS-CLKESCMRK-ACC# (501)
La3.3A	RPP-CCTYD---GS-CLKESCKRK-ACC# (502)
$\mu$ -GIIIA	-RD-CCTOO---KK-CKDRQCKOQ-RCCA# (503)
$\mu$ -GIIIB	-RD-CCTOO---RK-CKDRRCKOM-KCCA# (504)
30 $\mu$ -GIIIC	-RD-CCTOO---KK-CKDRRCKOL-KCCA# (505)
$\mu$ -PIIIA	ZRL-CCGFO---KS-CKRSRQCKOH-RCC# (506)
$\mu$ -PIIIA	-RD-CCTPP---KK-CKDRQCKPQ-RCCA# (507)
M3.1	RGG-CCTPP---RK-CKDRACKPA-RCCGP# (508)
Mr3.1	ZK--CCTGK---KGCSGKACKNL-KCCS# (509)
Nb3.2 (F582)	RGG-CCTPP---KK-CKDRACKPA-RCCGP# (510)
35 Pr3.1 (Z500)	-RG-CCTPP---RK-CKDRACKPA-RCCGP# (511)
Pr3.2 (Z501)	LOS-CCSLN---LRLCOVOACKRN-OCCT# (512)
R3.1	ZQR-CCTVK---RICOVOACRSK-OCCKS^ (513)
R3.2	RGG-CCTPP---RK-CKDRACKPA-RCCGP# (514)
R3.3	ZK--CCTGK---KGCSGKACKNL-KCCS# (515)
40 Sm3.1	H-G-CCKGO---EG-CSSRECROQ-HCC# (516)
T3.1	H-G-CCEGP---KG-CSSRECRPQ-HCC# (517)
T3.2 (Y088)	LPS-CCDFE---RLCVVPACIRH-QCCT# (518)
Wi3.1 (M548)	

Type 5

Om3.2

CCKYGWTCLLGCTPCDC^ (519)

Om3 . 4

CCRYGWTWCWLGCTPCGC^ (520)

Type 6

5

S3 . 2 (F352)

Z-NCCNGG-CSSKWCRDHARCC# (432)

## EXAMPLE 3

Effect of Intrathecal Administration of  $\mu$ -Conopeptides

10 [0087] Male C57 black mice (20-25g) are obtained from Charles River Laboratories. These mice and the animals are housed in a temperature controlled ( $23^\circ \pm 3^\circ$  C) room with a 12 hour light-dark cycle with free access to food and water. All animals are euthanized in accordance with Public Health Service policies on the humane care of laboratory animals.

15 [0088] Intrathecal (it) drug injections are performed as described (Hylden and Wilcox, 1980). A  $\mu$ -conopeptide or vehicle is administered in a volume of 5  $\mu$ l. Duration of hind-limb paralysis is assessed. This experiment reveals that injection of  $\mu$ -conopeptides into the intrathecal space of C57 black mice produced a paralysis of the animal. The animals in this experiment recovered fully.

20

## EXAMPLE 4

Effect of  $\mu$ -Conopeptides as a Local Anesthetic

25 [0089] Male Hartley guinea pigs (retired breeders) are obtained from Charles River Laboratories. The local anesthetic test is performed essentially as described (Bulbring and Wajda, 1945). On the day prior to test day, a patch on the back of the guinea pig is denuded of hair, first by shaving with electric clippers and subsequently with depilatory cream (Nair®). Depilatory cream is applied for five minutes and removed with a warm washcloth. The guinea pigs are dried and returned to their cages. On the following day, intradermal injections (0.1 ml vols) of lidocaine, bupivacaine, a  $\mu$ -conopeptide or vehicle (0.5% cyclodextran) are made into the denuded patch. The injection produced a raised wheal on the surface of the skin which is circled with a felt-tipped pen. Typically, four injections are made on the back of each guinea pig. In some cases, guinea pigs are reused following at least one week of recovery and injecting into an unused portion of the skin. The stimulus consists of mild pin pricks (not hard enough to break the skin) with a 26G needle. The response is a localized skin twitch caused by contraction

of cutaneous muscles. A unit test consisted of six uniform pin pricks, 3-5 seconds apart, within the injected area. Unit scores range from 0 (complete anesthesia) to 6 (no anesthesia). For potency experiments, the unit test is repeated at each site at five minute intervals for 30 minutes, and unit test scores summed (with 36 representing no anesthesia to 0 representing complete anesthesia). For duration experiments, unit tests are performed as described over the course of several hours to days.

[0090]  $\mu$ -Conopeptides of the present invention produce a potent and long lasting local anesthetic effect in the intracutaneous wheal test in the guinea pig. As expected, bupivacaine has a slightly longer duration than lidocaine, consistent with clinical observations.

10

#### EXAMPLE 5

##### Muscle Relaxant Effect of $\mu$ -Conopeptides in Anesthetized Monkeys

[0091]  $\mu$ -Conopeptides are dissolved 0.9 percent saline at a concentration of 2 mg/ml. Rhesus monkeys are anesthetized with halothane, nitrous oxide and oxygen. The maintenance concentration of halothane is 1.0%. Arterial and venous catheters are placed in the femoral vessels for drug administration and recording of the arterial pressure. Controlled ventilation is accomplished via an endotrachael tube. Twitch and tetanic contractions of the tibialis anterior muscle are elicited indirectly via the sciatic nerve. Recordings of arterial pressure electrocardiogram (lead I), heart rate, and muscle function are made simultaneously. Four to six animals received each listed compound. Four additional animals received succinylcholine chloride or d-tubocurarine chloride as controls. It is seen that the tested  $\mu$ -conopeptides generally provide similar or better results than those seen for succinylcholine chloride or d-tubocurarine chloride.

25

#### EXAMPLE 6

##### In vivo Activity of $\mu$ -Conopeptides in Pain Models

[0092] The anti-pain activity of  $\mu$ -conopeptides is shown in several animal models. These models include the nerve injury model (Chaplan, et al., 1997), the nociceptive response to s.c. formalin injection in rats (Codene, 1993) and an NMDA-induced persistent pain model (Liu, et al., 1997). In each of these models it is seen that the  $\mu$ -conopeptides and  $\mu$ -conopeptides derivatives have analgesic properties.

[0093] More specifically, this study evaluates the effect of intrathecal administration of  $\mu$ -conopeptides in mice models of nociceptive and neuropathic pain. For nociceptive pain, the effect of the  $\mu$ -conopeptides is studied in two different tests of inflammatory pain. The first is the formalin test, ideal because it produces a relatively short-lived, but reliable pain behavior that is readily quantified. There are two phases of pain behavior, the second of which is presumed to result largely from formalin-evoked inflammation of the hind paw. A  $\mu$ -conopeptide is administered 10 minutes prior to injection of formalin. The number of flinches and/or the duration of licking produced by the injection is monitored. Since the first phase is presumed to be due to direct activation of primary afferents, and thus less dependent on long term changes in the spinal cord,  $\mu$ -conopeptides are presumed to have greatest effect on the magnitude of pain behavior in the second phase.

[0094] The mechanical and thermal thresholds in animals that received an injection of complete Freund's adjuvant into the hind paw are also studied. This produces a localized inflammation including swelling of the hind paw and a profound decrease in mechanical and thermal thresholds, that are detected within 24 hours after injection. The changes in thresholds in rats that receive  $\mu$ -conopeptides are compared with those of rats that receive vehicle intrathecal injections.

[0095] An important issue is whether the drugs are effective when administered after the pain model has been established, or whether they are effective only if used as a pretreatment. Clearly, the clinical need is for drugs that are effective after the pain has developed. To address this issue, animals are studied in which  $\mu$ -conopeptides are administered repeatedly, after the inflammation (CFA) or nerve injury has been established. In these experiments, a  $\mu$ -conopeptide is injected daily by the intrathecal (i.t.) route. The mechanical and thermal thresholds (measured, respectively, with von Frey hairs in freely moving animals and with the Hargreave's test, also in freely moving animals) are repeated for a 2 to 4 week period after the injury is induced and the changes in pain measured monitored over time.

#### EXAMPLE 7

##### Effect of $\mu$ -Conotoxins in a Pain Model

[0096] Analgesic activity of  $\mu$ -conotoxins is also tested in pain models as follows.

[0097] Persistent pain (formalin test). Intrathecal (i.t) drug injections are performed as described by Hylden and Wilcox (1980). An  $\mu$ -conopeptide or vehicle is administered in a

volume of 5  $\mu$ l. Fifteen minutes after the i.t. injection, the right hindpaw is injected with 20  $\mu$ l of 5% formalin. Animals are placed in clear plexiglass cylinders backed by mirrors to facilitate observation. Animals are closely observed for 2 minutes per 5 minute period, and the amount of time the animal spent licking the injected paw is recorded in this manner for a total of 45-50 minutes. Results are expressed as licking time in seconds per five minutes. At the end of the experiment, all animals are placed on an accelerating rotarod and the latency to first fall was recorded.  $\mu$ -Conopeptides are found to be active in this model which is predictive of efficacy for treating neuropathic pain.

[0098] Acute pain (tail-flick). A  $\mu$ -conopeptide or saline is administered intrathecally (i.t.) according to the method of Hylden and Wilcox (1980) in a constant volume of 5  $\mu$ l. Mice are gently wrapped in a towel with the tail exposed. At various time-points following the i.t. injection, the tail is dipped in a water bath maintained at 54° C. and the time to a vigorous tail withdrawal is recorded. If there is no withdrawal by 8 seconds, the tail is removed to avoid tissue damage.

[0099] Neuropathic pain. The partial sciatic nerve ligation model is used to assess the efficacy of  $\mu$ -conopeptides in neuropathic pain. Nerve injury is produced according to the methods of Malmberg and Basbaum (1998). Animals are anesthetized with a ketamine/xylazine solution, the sciatic nerve is exposed and tightly ligated with 8-0 silk suture around 1/3 to  $\frac{1}{2}$  of the nerve. In sham-operated mice the nerve is exposed, but not ligated. Animals are allowed to recover for at least 1 week before testing is performed. On the testing day, mice are placed in plexiglass cylinders on a wire mesh frame and allowed to habituate for at least 60 minutes. Mechanical allodynia is assessed with calibrated von Frey filaments using the up-down method as described by Chaplan et al. (1994), and the 50% withdrawal threshold is calculated. Animals that did not respond to any of the filaments in the series are assigned a maximal value of 3.6 grams, which is the filament that typically lifted the hindlimb without bending, and corresponds to approximately 1/10 the animal's body weight.

[0100] The data obtained demonstrate that  $\mu$ -conopeptides have potent analgesic properties in three commonly used models of pain: acute, persistent/inflammatory and neuropathic pain models.

## EXAMPLE 8

Activity of  $\mu$ -Conopeptide S3.2 on Neuronal Sodium Channels

[0101]  $\mu$ -Conopeptide S3.2 was tested for activity on sodium channels as follows. S3.2 was administered to mice by intracerebroventricular (ICV) injection. Administration of S3.2 in 5 this manner caused mice to show a spectrum of activity that is characteristic of all sodium channel blockers, including rapid loss of righting reflex, coma-like inactivity and spastic uncontrolled limb movement. Following intrathecal (it) administration to mice, S3.2 causes rapid hindlimb paralysis that spreads to include the entire body over a course of 10-20 minutes followed by death, presumably due to respiratory paralysis. However, unlike classic  $\mu$ -conopeptides, S3.2 has no significant activity following intravenous administration (iv) to mice. 10 Classic  $\mu$ -conopeptides, such as GI<sub>IIIA</sub> and PI<sub>IIIA</sub>, cause rapid paralysis and death following iv administration, indicating their activity at skeletal muscle sodium channels. To confirm the selectivity of S3.2, 80 nmol was administered iv to rats. The effect of S3.2 was measured on skeletal muscle contraction, blood pressure and heart rate. S3.2 was found to have no effect on 15 any of these parameters. Controls were performed using classical  $\mu$ -conopeptides, including Sm3.1, Sm3.3 and Bu3.1 described herein, also administered iv at 80 nmol. These control peptides caused a dramatic decrease in skeletal muscle contractility, as well as a significant drop in systemic blood pressure. Thus,  $\mu$ -conopeptide S3.2 surprisingly is selective for neuronal sodium channels. The most obvious difference between the S3.2 sequence and the sequences of 20 these other peptides is a shortened first loop (the first loop between cysteine residues) which lacks a charged amino acid.

[0102] It will be appreciated that the methods and compositions of the instant invention can be incorporated in the form of a variety of embodiments, only a few of which are disclosed 25 herein. It will be apparent to the artisan that other embodiments exist and do not depart from the spirit of the invention. Thus, the described embodiments are illustrative and should not be construed as restrictive.

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U.S. Patent No. 5,670,622.

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Published PCT Application WO 95/01203.

- Published PCT Application WO 95/05452.
- Published PCT Application WO 96/02286.
- Published PCT Application WO 96/02646.
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WHAT IS CLAIMED IS:

1. An isolated peptide selected from the group consisting of:
  - (a) a peptide set forth in Table 1 or Table 2 and
  - 5 (b) a derivative of the peptide in (a).
2. The isolated peptide of claim 1, wherein Xaa<sub>1</sub> is Glu, Xaa<sub>2</sub> is pyro-Glu, Xaa<sub>4</sub> is Trp and Xaa<sub>5</sub> is Tyr.
- 10 3. The derivative of the peptide of claim 1, in which the Arg residues may be substituted by Lys, ornithine, homoarginine, nor-Lys, N-methyl-Lys, N,N-dimethyl-Lys, N,N,N-trimethyl-Lys or any synthetic basic amino acid; the Lys residues may be substituted by Arg, ornithine, homoarginine, nor-Lys, or any synthetic basic amino acid; the Tyr residues may be substituted with meta-Tyr, ortho-Tyr, nor-Tyr, mono-halo-Tyr, di-halo-Tyr, O-sulpho-Tyr, O-phospho-Tyr, nitro-Tyr or any synthetic hydroxy containing amino acid; the Ser residues may be substituted with Thr or any synthetic hydroxylated amino acid; the Thr residues may be substituted with Ser or any synthetic hydroxylated amino acid; the Phe residues may be substituted with any synthetic aromatic amino acid; the Trp residues may be substituted with Trp (D), neo-Trp, halo-Trp (D or L) or any aromatic 15 synthetic amino acid; the Asn, Ser, Thr or Hyp residues may be glycosylated; the Tyr residues may also be substituted with the 3-hydroxyl or 2-hydroxyl isomers (meta-Tyr or ortho-Tyr, respectively) and corresponding O-sulpho- and O-phospho-derivatives; the acidic amino acid residues may be substituted with any synthetic acidic amino acid, e.g., tetrazolyl derivatives of Gly and Ala; the aliphatic amino acids may be substituted by 20 synthetic derivatives bearing non-natural aliphatic branched or linear side chains C<sub>n</sub>H<sub>2n+2</sub> up to and including n=8; the Met residues may be substituted by Nle; the Cys residues may be in D or L configuration and may optionally be substituted with homocysteine (D or L); pairs of Cys residues may be replaced pairwise with isoteric lactam or ester-thioether replacements, such as Ser/(Glu or Asp), Lys/(Glu or Asp), Cys/Glu (or Asp) or 25 Cys/Ala combinations; and the peptide may be radioiodinated or radiotriated.

4. A substantially pure  $\mu$ -conotoxin peptide derivative comprising a permutant of the peptide of claim 1,2 or 3.
5. A substantially pure  $\mu$ -conotoxin peptide derivative comprising the peptide or peptide derivative of claim 1, 2 or 3 modified to contain an O-glycan, an S-glycan or an N-glycan.
6. A substantially pure  $\mu$ -conotoxin peptide derivative comprising the peptide derivative of claim 4 modified to contain an O-glycan, an S-glycan or an N-glycan.

10

7. An isolated nucleic acid encoding a  $\mu$ -conopeptide propeptide having an amino acid sequence set forth in Table 1.
8. The isolated nucleic acid of claim 7, wherein the nucleic acid comprises a nucleotide sequence set forth in Table 1.
- 15
9. An isolated  $\mu$ -conopeptide propeptide having an amino acid sequence set forth in Table 1.

20

10. A method for treating or preventing disorders associated with voltage gated ion channel disorders in which comprises administering to a patient in need thereof a therapeutically effective amount of a peptide of claim 1 or a pharmaceutically acceptable salt thereof.
11. The method of claim 10, wherein said disorder is a neurologic disorder.

25

12. The method of claim 11, wherein said neurologic disorder is Amyotrophic Lateral Sclerosis.
13. The method of claim 11, wherein said neurologic disorder is head trauma.

30

14. The method of claim 11, wherein said neurologic disorder is epilepsy.

15. The method of claim 11, wherein said neurologic disorder is a neurotoxic injury associated with conditions of hypoxia, anoxia or ischemia.
16. The method of claim 15, wherein said neurotoxic injury is associated with stroke, cerebrovascular accident, brain or spinal cord trauma, myocardial infarct, physical trauma, drownings, suffocation, perinatal asphyxia, or hypoglycemic events.
17. The method of claim 10, wherein said disorder is pain.
- 10 18. The method of claim 17, wherein said pain is migraine, acute pain, persistent pain, chronic pain, neuropathic pain or nociceptive pain.
19. The method of claim 18, wherein the pain is trigeminal neuralgia, diabetic neuropathy, post-herpetic neuralgia, neuroma pain, phantom limb pain.
- 15 20. The method of claim 17, wherein said pain is burn pain.
21. The method of claim 10, wherein said disorder is a neuromuscular disorder.
- 20 22. The method of claim 21, wherein said neuromuscular disorder is myofacial pain syndrome, chronic muscle spasm, dystonias or spasticity.
23. A method for providing musculoskeletal relaxation in a patient undergoing a surgical procedure requiring anesthesia which comprises administering to a patient in need thereof a therapeutically effective amount of a peptide of claim 1 or a pharmaceutically acceptable salt thereof.
- 25 24. A method of alleviating pain which comprises administering to a mammal that is either exhibiting pain or is about to be subjected to a pain-causing event a pain-alleviating amount of a peptide of claim 1 or a pharmaceutically acceptable salt thereof.
- 30 25. The method of claim 24, wherein the peptide is administered as a local anesthetic.

26. The method of claim 24, wherein the peptide is administered as an ocular anesthetic.
27. A method for characterizing a pore occlusion site on a sodium channel subtype  
5 comprising determining the affinity of said site for a peptide of claim 1.
28. The method of claim 27, wherein said sodium channel subtype is a neuronal sodium channel subtype and said peptide is  $\mu$ -conopeptide S3.2 comprising an amino acid sequence as set forth in SEQ ID NO:211 or SEQ ID NO:432.  
10
29. A method for screening a small molecule library to identify a small molecule which is a selective blocking agent of a sodium channel subtype comprising (a) measuring the blocking activity of a small molecule on said sodium channel subtype, (b) measuring the blocking activity of a peptide of claim 1 on said sodium channel subtype and (c) comparing the blocking activity of said small molecule with the blocking activity of said peptide.  
15
30. The method of claim 29, wherein said sodium channel subtype is a neuronal sodium channel subtype and said peptide is  $\mu$ -conopeptide S3.2 comprising an amino acid sequence as set forth in SEQ ID NO:211 or SEQ ID NO:432.  
20
31. A method for screening a small molecule library to identify a small molecule which is a selective blocking agent of a sodium channel subtype comprising (a) measuring the binding affinity of a small molecule on said sodium channel subtype, (b) measuring the binding affinity of a peptide of claim 1 on said sodium channel subtype and (c) comparing the binding affinity of said small molecule with the binding affinity of said peptide.  
25
32. The method of claim 31, wherein said peptide is radiolabeled.  
30

33. The method of claim 31, wherein said sodium channel subtype is a neuronal sodium channel subtype and said peptide is  $\mu$ -conopeptide S3.2 comprising an amino acid sequence as set forth in SEQ ID NO:211 or SEQ ID NO:432.
- 5 34. The method of claim 33, wherein said peptide is radiolabeled.
35. A method for screening a small molecule library to identify a small molecule which is a selective blocking agent of a sodium channel subtype comprising (a) allowing a peptide of claim 1 to bind to a sodium channel subtype, (b) adding a small molecule and (c) measuring the amount of displacement of said peptide on said sodium channel subtype by said small molecule.
- 10 36. The method of claim 35, wherein said peptide is radiolabeled.
- 15 37. The method of claim 35, wherein said sodium channel subtype is a neuronal sodium channel subtype and said peptide is  $\mu$ -conopeptide S3.2 comprising an amino acid sequence as set forth in SEQ ID NO:211 or SEQ ID NO:432.
- 20 38. The method of claim 37, wherein said peptide is radiolabeled.
39. A method for screening a small molecule library to identify a small molecule which is a selective blocking agent of a sodium channel subtype comprising (a) allowing a small molecule to bind to a sodium channel subtype, (b) adding a peptide of claim 1 and (c) measuring the amount of displacement of said small molecule on said sodium channel subtype by said small peptide.
- 25 40. The method of claim 39, wherein said sodium channel subtype is a neuronal sodium channel subtype and said peptide is  $\mu$ -conopeptide S3.2 comprising an amino acid sequence as set forth in SEQ ID NO:211 or SEQ ID NO:432.
- 30 41. A method of identifying compounds that mimic the therapeutic activity of a  $\mu$ -conotoxin, comprising the steps of: (a) conducting a biological assay on a test compound to

determine the therapeutic activity; and (b) comparing the results obtained from the biological assay of the test compound to the results obtained from the biological assay of a  $\mu$ -conotoxin, wherein said  $\mu$ -conotoxin is a peptide of claim 1.

5 42. The method of claim 41, wherein said  $\mu$ -conotoxin is S3.2 comprising an amino acid set forth in SEQ ID NO:211 or SEQ IN NO:432.

## SEQUENCE LISTING

<110> University of Utah Research Foundation  
CogneRx, Inc.  
Olivera, Baldomero M.  
McIntosh, J. Michael  
Garrett, James E.  
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Shon, Ki-Joon  
Jacobsen, Richard  
Jones, Robert M.  
Cartier, G. Edward  
Shen, Greg S.  
Wagstaff, John D.

<120> Mu-Conopeptides

<130> 2314-242

<150> US 60/219,619  
<151> 2000-07-21

<150> US 60/245,157  
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agagcgtatg caggacgact ttataactga gcatcatccc ctgtttgatc ctgtcaaacg 180  
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Ala Glu Arg Met Gln Asp Asp Phe Ile Thr Glu His His Pro Leu Phe

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45

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 gtgttgcggc gaaggccgga aatgccccag ctatccaga aacagtcaga tttgtcattg      240  
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Glu Glu Arg Met Gln Asp Asp Ile Ser Ser Glu Gln His Pro Leu Phe  
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Arg Pro Ala Glu Arg Met Gln Asp Asp Ile Ser Ser Glu Gln His Arg  
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 <213> Conus betulinus

<400> 29  
 Met Met Phe Lys Leu Gly Val Leu Leu Thr Ile Tyr Met Leu Leu Phe  
 1 5 10 15

Pro Phe Thr Ala Leu Pro Leu Asp Gly Asp Gln Pro Ala Asp Gln Pro  
 20 25 30

Leu Glu Arg Met Gln Tyr Asp Met Leu Arg Ala Val Asn Pro Trp Phe  
 35 40 45

Asp Pro Val Lys Arg Cys Cys Ser Arg Asn Cys Ala Val Cys Ile Pro  
 50 55 60

Cys Cys Pro Asn Trp Pro Ala  
 65 70

<210> 30  
 <211> 18  
 <212> PRT  
 <213> Conus betulinus

<220>  
 <221> PEPTIDE  
 <222> (1)...(18)  
 <223> Xaa at residue 11, 14 and 17 is Pro or Hyp; Xaa at residue 16 is  
 Trp or bromo-Tr

<400> 30  
 Cys Cys Ser Arg Asn Cys Ala Val Cys Ile Xaa Cys Cys Xaa Asn Xaa  
 1 5 10 15

Xaa Ala

<210> 31  
 <211> 325  
 <212> DNA  
 <213> Conus bullatus

<400> 31  
 caagaaggat cgatagcagt tcatgtatgc taaaactggga gtcttggat ccatctgtct 60  
 gcttctgttt cccctttttt ctcttccgca ggatggagat caacctgcag accgacacctgc 120  
 agagcgtatg caggacgaca tttcatctga gcagaattcc ttgcttggaa agagagttac 180  
 tgacaggtgc tgcaaaggga agagggatgg cggcagatgg tgcagagatc actcgcgttg 240  
 ttgcggtcga cgataagctg ttgatgacca gcttttttat cacggctaca tcaagtgtct 300  
 agtgaataag taaaatgatt gcagt 325  
 <210> 32

<211> 77  
 <212> PRT  
 <213> Conus bullatus

<400> 32  
 Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15

Pro Leu Phe Ala Leu Pro Gln Asp Gly Asp Gln Pro Ala Asp Arg Pro  
 20 25 30

Ala Glu Arg Met Gln Asp Asp Ile Ser Ser Glu Gln Asn Ser Leu Leu  
 35 40 45

Glu Lys Arg Val Thr Asp Arg Cys Cys Lys Gly Lys Arg Glu Cys Gly  
 50 55 60

Arg Trp Cys Arg Asp His Ser Arg Cys Cys Gly Arg Arg  
 65 70 75

<210> 33  
 <211> 23  
 <212> PRT  
 <213> Conus bullatus

<220>  
 <221> PEPTIDE  
 <222> (1)..(23)  
 <223> Xaa at residue 11 is Glu or gamma-carboxy Glu; Xaa at residue 15  
 is Trp or bromo-Tr

<400> 33  
 Val Thr Asp Arg Cys Cys Lys Gly Lys Arg Xaa Cys Gly Arg Xaa Cys  
 1 5 10 15

Arg Asp His Ser Arg Cys Cys  
 20

<210> 34  
 <211> 326  
 <212> DNA  
 <213> Conus bullatus

<400> 34  
 caagaggat cgatagcagt tcatgatgtc taaaactggga gtcttggta ccatctgtct 60  
 gcttctgttt cccctttttg ctcttcggca ggatggagat caacctgcag accgacacctgc 120  
 agagcgtatg caggtatgaca tttcatctga gcagaatccc ttgcttgaga agagagttgg 180  
 tgacagggtgc tgcaaaggga agagggggtg cggcagatgg tgcagagatc actcacgttg 240  
 ttgcggtcga cgataacgtg ttgatgacca gctttgttat cacggctaca tcaagtgtct 300  
 tagtgattaa gtaaaacgtat tgcagt 326

<210> 35  
 <211> 77  
 <212> PRT  
 <213> Conus bullatus

<400> 35  
 Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15

Pro Leu Phe Ala Leu Arg Gln Asp Gly Asp Gln Pro Ala Asp Arg Pro  
20 25 30

Ala Glu Arg Met Gln Asp Asp Ile Ser Ser Glu Gln Asn Pro Leu Leu  
35 40 45

Glu Lys Arg Val Gly Asp Arg Cys Cys Lys Gly Lys Arg Gly Cys Gly  
50 55 60

Arg Trp Cys Arg Asp His Ser Arg Cys Cys Gly Arg Arg  
65 70 75

<210> 36

<211> 23

<212> PRT

<213> Conus bullatus

<220>

<221> PEPTIDE

<222> (1)..(23)

<223> Xaa at residue 15 is Trp or bromo-Trp

<400> 36

Val Gly Asp Arg Cys Cys Lys Gly Lys Arg Gly Cys Gly Arg Xaa Cys  
1 5 10 15

Arg Asp His Ser Arg Cys Cys  
20

<210> 37

<211> 331

<212> DNA

<213> Conus bullatus

<400> 37

caagaaggat cgatacgagt tcatgatgtc taaaactggga gtcttgttga ccatctgtct 60

gcttctgtt ccccttttg ctcttccgca ggtggagat caacctgcag accgacactgc 120

agagcgtatg caggacgaca ttcatctga gcagaatccc ttgcttgaga agagagttgg 180

tgaaagggtgc tgcaaaaacg ggaagagggg gtgcggcaga tgggcagag atcactcagc 240

ttgttgcgtt cgacgataac gtgttcatgtca ccgaggctt cgatcatcagc gctacatcaa 300

gtgtcttagtg aataagtaaa acgattgcag t 331

<210> 38

<211> 78

<212> PRT

<213> Conus bullatus

<400> 38

Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
1 5 10 15

Pro Leu Phe Ala Leu Pro Gln Asp Gly Asp Gln Pro Ala Asp Arg Pro  
20 25 30

Ala Glu Arg Met Gln Asp Asp Ile Ser Ser Glu Gln Asn Pro Leu Leu  
35 40 45

Glu Lys Arg Val Gly Glu Arg Cys Cys Lys Asn Gly Lys Arg Gly Cys  
50 55 60

Gly Arg Trp Cys Arg Asp His Ser Arg Cys Cys Gly Arg Arg

65	70	75	
<210> 39 <211> 24 <212> PRT <213> Conus bullatus			
<220> <221> PEPTIDE <222> (1)..(24) <223> Xaa at residue 3 is Glu or gamma-carboxy Glu; Xaa at residue 16 is Trp or bromo-Tr			
<400> 39 Val Gly Xaa Arg Cys Cys Lys Asn Gly Lys Arg Gly Cys Gly Arg Xaa 1 5 10 15			
Cys Arg Asp His Ser Arg Cys Cys 20			
<210> 40 <211> 337 <212> DNA <213> Conus bullatus			
<400> 40 caagaggat cgatacgagt tcatgtatgtc taaaactggga gtcttggta ccatctgtct 60 gcttctgttt ccccttttg ctcttccgca ggacggagat caacctgcag accgacactgc 120 agagcgtatg caggacgacc tttcatctga gcagcatccc ttgtttgaga agagaattgt 180 tgacagggtgc tgcaacaaag ggaacgggaa gaggggggtgc agcagatggc gcagagatca 240 ctcacgttgt tgccgtcgac gatgaactgt tgatgaccga ggctttggtt atcacggcta 300 catcaagtgt ctatgtata agtaaaacga ttgcagt 337			
<210> 41 <211> 80 <212> PRT <213> Conus bullatus			
<400> 41 Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe 1 5 10 15			
Pro Leu Phe Ala Leu Pro Gln Asp Gly Asp Gln Pro Ala Asp Arg Pro 20 25 30			
Ala Glu Arg Met Gln Asp Asp Leu Ser Ser Glu Gln His Pro Leu Phe 35 40 45			
Glu Lys Arg Ile Val Asp Arg Cys Cys Asn Lys Gly Asn Gly Lys Arg 50 55 60			
Gly Cys Ser Arg Trp Cys Arg Asp His Ser Arg Cys Cys Gly Arg Arg 65 70 75 80			
<210> 42 <211> 26 <212> PRT <213> Conus bullatus			
<220>			

&lt;221&gt; PEPTIDE

&lt;222&gt; (1)..(26)

&lt;223&gt; Xaa at residue 18 is Trp or bromo-Trp

&lt;400&gt; 42

Ile Val Asp Arg Cys Cys Asn Lys Gly Asn Gly Lys Arg Gly Cys Ser  
1 5 10 15Arg Xaa Cys Arg Asp His Ser Arg Cys Cys  
20 25

&lt;210&gt; 43

&lt;211&gt; 337

&lt;212&gt; DNA

&lt;213&gt; Conus bullatus

&lt;400&gt; 43

caagaaggat cgatacgagt tcatgatgtc taaaactggga gtcttggta ccatctgtct 60

gcttctgttt cccctttttg ctcttccgca ggatggagat caacctgcag accgacactgc 120

tgagcgtatg caggacgaca tttcatctga gcggaatccc ttgtttgaga agagcggtgg 180

tttatattgc tgccgaccca aacccaacgg gcagatgtatg tgcgacagat ggtgcgaaaa 240

aaactcacgt tggcggtc gacgataatg tggatgac cagctttgtt atcaaggcta 300

catcaagttat ctatgtataa agtaaaaacga ttgcagt 337

&lt;210&gt; 44

&lt;211&gt; 77

&lt;212&gt; PRT

&lt;213&gt; Conus bullatus

&lt;400&gt; 44

Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
1 5 10 15Pro Leu Phe Ala Leu Pro Gln Asp Gly Asp Gln Pro Ala Asp Arg Pro  
20 25 30Ala Glu Arg Met Gln Asp Asp Ile Ser Ser Asn Pro Leu Phe Glu Lys  
35 40 45Ser Val Gly Cys Cys Arg Pro Lys Pro Asn Gly Gln Met Met Cys Asp  
50 55 60Arg Trp Cys Glu Lys Asn Ser Arg Cys Cys Gly Arg Arg  
65 70 75

&lt;210&gt; 45

&lt;211&gt; 27

&lt;212&gt; PRT

&lt;213&gt; Conus bullatus

&lt;220&gt;

&lt;221&gt; PEPTIDE

&lt;222&gt; (1)..(27)

&lt;223&gt; Xaa at residue 21 is Glu or gamma-carboxy Glu; Xaa at residue 8 and 10 is Pro or Hyp; Xaa at residue 19 is Trp or bromo-Trp; Xaa at residue 4 is Tyr, 125I-Tyr, mono-iodo-Tyr, di-iodo-Tyr, O-sulpho-Tyr or O-phospho-Ty

&lt;400&gt; 45

Val Gly Leu Xaa Cys Cys Arg Xaa Lys Xaa Asn Gly Gln Met Met Cys

14

1 5 10 15

Asp Arg Xaa Cys Xaa Lys Asn Ser Arg Cys Cys  
20 25

<210> 46  
<211> 323  
<212> DNA  
<213> Conus bullatus  
<400> 46  
caagaaggat cgatacgagt tcatgatgtc taaactggga gttttgttga ccatctgtct 60  
gcttctgttt ccccttaactg ctcttccgat ggatggagat caatctgttag accgacactgc 120  
agaacgtatg caggacgacc tttcatctga gcagcatccc ttgtttgttc agaaaagaag 180  
gtgttgcggc gaaggcttga catgccccag atattggaaa aacagtcaga tttgtgcttg 240  
ttgttaaatg acaacgtgtc gatgaccaac ttccgtatca cgactacgcc aagtgtctaa 300  
tgaataagta aaacgattgc agt 323

<210> 47  
<211> 74  
<212> PRT  
<213> Conus bullatus

<400> 47  
Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
1 5 10 15

Pro Leu Thr Ala Leu Pro Met Asp Gly Asp Gln Ser Val Asp Arg Pro  
20 25 30

Ala Glu Arg Met Gln Asp Asp Leu Ser Ser Glu Gln His Pro Leu Phe  
35 40 45

Val Gln Lys Arg Arg Cys Cys Gly Glu Gly Leu Thr Cys Pro Arg Tyr  
50 55 60

Trp Lys Asn Ser Gln Ile Cys Ala Cys Cys  
65 70

<210> 48  
<211> 22  
<212> PRT  
<213> Conus bullatus

<220>  
<221> PEPTIDE  
<222> (1)..(22)  
<223> Xaa at residue 5 is Glu or gamma-carboxy Glu; Xaa at residue 10 is Pro or Hyp; Xaa at residue 13 is Trp or bromo-Trp; Xaa at residue 12 is Tyr, 125I-Tyr, mono-iodo-Tyr, di-iodo-Tyr, O-sulpho-Tyr or O-phospho-Tyr

<400> 48  
Arg Cys Cys Gly Xaa Gly Leu Thr Cys Xaa Arg Xaa Xaa Lys Asn Ser  
1 5 10 15

Gln Ile Cys Ala Cys Cys  
20

<210> 49  
<211> 322

&lt;212&gt; DNA

&lt;213&gt; Conus bullatus

&lt;400&gt; 49

caaggaggat	cgatacgagt	tcatgatgtc	taaactggga	gtcttgttga	ccatctgtct	60
gcttctttt	ccccctttt	ctcttccgca	ggatggagat	caacctgcag	accgaccctgc	120
tgagcgtatg caggacgaca tttcatctga gcaggatccc ttgtttgttc agaaaagaag						180
gtgttgcggc gaaggcttga catgccccag atattggaaa aacagtcaga tttgtgcttg						240
ttgttaaatg acaacgtgtg atgaccaaact tcggtatcac gactacgcca agtgtctaat						300
gaataagtaa aacgattgca gt						322

&lt;210&gt; 50

&lt;211&gt; 74

&lt;212&gt; PRT

&lt;213&gt; Conus bullatus

&lt;400&gt; 50

Met	Met	Ser	Lys	Leu	Gly	Val	Leu	Leu	Thr	Ile	Cys	Leu	Leu	Phe
1				5				10				15		

Pro	Leu	Phe	Ala	Leu	Pro	Gln	Asp	Gly	Asp	Gln	Pro	Ala	Asp	Arg	Pro
				20			25				30				

Ala	Glu	Arg	Met	Gln	Asp	Asp	Ile	Ser	Ser	Glu	Gln	Asp	Pro	Leu	Phe
			35			40				45					

Val	Gln	Lys	Arg	Arg	Cys	Cys	Gly	Glu	Gly	Leu	Thr	Cys	Pro	Arg	Tyr
				50		55			60						

Trp	Lys	Asn	Ser	Gln	Ile	Cys	Ala	Cys	Cys
65					70				

&lt;210&gt; 51

&lt;211&gt; 22

&lt;212&gt; PRT

&lt;213&gt; Conus bullatus

&lt;220&gt;

&lt;221&gt; PEPTIDE

&lt;222&gt; (1)..(22)

<223> Xaa at residue 5 is Glu or gamma-carboxy Glu; Xaa at residue 10 is Pro or Hyp; Xaa at residue 13 is Trp or bromo-Trp; Xaa at residue 12 is Tyr, 125I-Tyr, mono-iodo-Tyr, di-iodo-Tyr, O-sulpho-Tyr or O-phospho-Ty

&lt;400&gt; 51

Arg	Cys	Cys	Gly	Xaa	Gly	Leu	Thr	Cys	Xaa	Arg	Xaa	Xaa	Lys	Asn	Ser
1				5			10			15					

Gln	Ile	Cys	Ala	Cys	Cys
			20		

&lt;210&gt; 52

&lt;211&gt; 238

&lt;212&gt; DNA

&lt;213&gt; Conus capitaneus

&lt;400&gt; 52

ggatccatga	tgtctaaact	gggagtcttg	gtgaccatct	gcctgcttct	gtttccctt	60
gctgctttc cactggatgg aaatcaacct gcagaccacc ctgcaaagcg tacgcaagat						120

gacagttcag ctgcctgtat caatacctgg attgatcatt cccattcttgc ctgcaggac 180  
 tgcggtaag attgtgttgg ttgtgccgg taacgtgttgc atgaccaact ttctcgag 238

<210> 53  
 <211> 70  
 <212> PRT  
 <213> *Conus capitaneus*

<400> 53  
 Gly Ser Met Met Ser Lys Leu Gly Val Leu Val Thr Ile Cys Leu Leu  
 1 5 10 15

Leu Phe Pro Leu Ala Ala Phe Pro Leu Asp Gly Asn Gln Pro Ala Asp  
 20 25 30

His Pro Ala Lys Arg Thr Gln Asp Asp Ser Ser Ala Ala Leu Ile Asn  
 35 40 45

Thr Trp Ile Asp His Ser His Ser Cys Cys Arg Asp Cys Gly Glu Asp  
 50 55 60

Cys Val Gly Cys Cys Arg  
 65 70

<210> 54  
 <211> 15  
 <212> PRT  
 <213> *Conus capitaneus*

<220>  
 <221> PEPTIDE  
 <222> (1)..(15)  
 <223> Xaa at residue 8 is Glu or gamma-carboxy Glu

<400> 54  
 Ser Cys Cys Arg Asp Cys Gly Xaa Asp Cys Val Gly Cys Cys Arg  
 1 5 10 15

<210> 55  
 <211> 323  
 <212> DNA  
 <213> *Conus characteristicus*

<400> 55  
 caagaggat cgatagcagt tcatgatgtc taaaactggga gtcttggta ccatctgtct 60  
 gcttctgttt ccccttactg ctcttccat ggatggagat caacctgcag accaacctgc 120  
 agatcgatg caggacgaca tttcatctga gcagtatccc ttgtttgata tgagaaaaag 180  
 gtgttgcggc cccggcgggtt catgccccgt atattcaga gacaattta ttgtgggtt 240  
 ttgttaatg acaacgtgtc gatgaccaac ttcattatca cgactacgcc aagtgtctaa 300  
 tgaataagta aaatgattgc agt 323

<210> 56  
 <211> 74  
 <212> PRT  
 <213> *Conus characteristicus*

<400> 56

Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15

Pro Leu Thr Ala Leu Pro Met Asp Gly Asp Gln Pro Ala Asp Gln Pro  
 20 25 30

Ala Asp Arg Met Gln Asp Asp Ile Ser Ser Glu Gln Tyr Pro Leu Phe  
 35 40 45

Asp Met Arg Lys Arg Cys Cys Gly Pro Gly Gly Ser Cys Pro Val Tyr  
 50 55 60

Phe Arg Asp Asn Phe Ile Cys Gly Cys Cys  
 65 70

<210> 57

<211> 21

<212> PRT

<213> Conus characteristicus

<220>

<221> PEPTIDE

<222> (1)..(21)

<223> Xaa at residue 4 and 9 is Pro or Hyp; Xaa at residue 11 is Tyr, 1  
 25I-Tyr, mono-iodo-Tyr, di-iodo-Tyr, O-sulpho-Tyr or O-phospho-Ty

<400> 57

Cys Cys Gly Xaa Gly Gly Ser Cys Xaa Val Xaa Phe Arg Asp Asn Phe  
 1 5 10 15

Ile Cys Gly Cys Cys  
 20

<210> 58

<211> 316

<212> DNA

<213> Conus characteristicus

<400> 58

caagaggat cgatagcagt tcatgatgtc taaaactggga gtcttggta ccatctgtct 60

gcttctgttt ccccttactg ctcttccgat ggtatggagat gaacctgcaa accgacctgt 120

cgagcgtatg caggacaaca tttcatctga gcagtatccc ttgtttgaga agagacgaga 180

ttgttgact cccggcaaga aatgcaaaga ccgacaatgc aaaccccaga gatgttgcgc 240

tggacgataa cgtgttcatg accaacttta tcacggctac gtcaagtgtt tagtgaataa 300

gtaaaatgtat tgcagt 316

<210> 59

<211> 75

<212> PRT

<213> Conus characteristicus

<400> 59

Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15

Pro Leu Thr Ala Leu Pro Met Asp Gly Asp Glu Pro Ala Asn Arg Pro  
 20 25 30

Val Glu Arg Met Gln Asp Asn Ile Ser Ser Glu Gln Tyr Pro Leu Phe

35

40

45

Glu Lys Arg Arg Asp Cys Cys Thr Pro Pro Lys Lys Cys Lys Asp Arg  
 50 55 60

Gln Cys Lys Pro Gln Arg Cys Cys Ala Gly Arg  
 65 70 75

&lt;210&gt; 60

&lt;211&gt; 22

&lt;212&gt; PRT

&lt;213&gt; Conus characteristicus

&lt;220&gt;

&lt;221&gt; PEPTIDE

&lt;222&gt; (1)..(22)

&lt;223&gt; Xaa at residue 6, 7 and 17 is Pro or Hyp

&lt;400&gt; 60

Arg Asp Cys Cys Thr Xaa Xaa Lys Lys Cys Lys Asp Arg Gln Cys Lys  
 1 5 10 15

Xaa Gln Arg Cys Cys Ala  
 20

&lt;210&gt; 61

&lt;211&gt; 314

&lt;212&gt; DNA

&lt;213&gt; Conus characteristicus

&lt;400&gt; 61

caagagggtt cgatagcagt tcattatgttc taaaactggga gtcttggatccatctgtct 60

gcttctgttt ccccttactg ctcttccact ggtggagat caacctgcag atcaatctgc 120

agagcgtacctt gcagagcgta cgcaggacga cattcagcag catccgttat atgatccgaa 180

aagaagggtt tgccgttatac catgccccga cagctgccac ggatcttgct gctataagt 240

ataacatgtt gatggccagc tttgttatca cggccacgtc aagtgtctta atgaataagt 300

aaaacgttgg cagt 314

&lt;210&gt; 62

&lt;211&gt; 72

&lt;212&gt; PRT

&lt;213&gt; Conus characteristicus

&lt;400&gt; 62

Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15

Pro Leu Thr Ala Leu Pro Leu Asp Gly Asp Gln Pro Ala Asp Gln Ser

20 25 30

Ala Glu Arg Pro Ala Glu Arg Thr Gln Asp Asp Ile Gln Gln His Pro

35 40 45

Leu Tyr Asp Pro Lys Arg Arg Cys Cys Arg Tyr Pro Cys Pro Asp Ser

50 55 60

Cys His Gly Ser Cys Cys Tyr Lys  
 65 70

&lt;210&gt; 63

&lt;211&gt; 18

&lt;212&gt; PRT

&lt;213&gt; Conus characteristicus

&lt;220&gt;

&lt;221&gt; PEPTIDE

&lt;222&gt; (1)..(18)

&lt;223&gt; Xaa at residue 6 and 8 is Pro or Hyp; Xaa at residue 5 and 17 is Tyr, 125I-Tyr, mono-iodo-Tyr, di-iodo-Tyr, O-sulpho-Tyr or O-phospho-Ty

&lt;400&gt; 63

Arg Cys Cys Arg Xaa Xaa Cys Xaa Asp Ser Cys His Gly Ser Cys Cys  
1 5 10 15

Xaa Lys

&lt;210&gt; 64

&lt;211&gt; 292

&lt;212&gt; DNA

&lt;213&gt; Conus characteristicus

&lt;400&gt; 64

caagaggat cgatagcagt tcatgatgtc taaaactggga gccttggta ccatctgtct 60  
acttctgtt tcccttactg ctgttccgct ggatggagat caacatgcag accaacacctc  
acagcgtctg caggaccgca ttccaactga agatcatccc ttatttgcac ccaacaaacg 120  
gtgttgcggcc cggtggcat gcaacatggg atgcaagcct tggtgtggat gaccagctt 180  
gttatcgccggttcatgaa gtgtcttaat gaataagtaa aatgattgca gt 240  
292

&lt;210&gt; 65

&lt;211&gt; 69

&lt;212&gt; PRT

&lt;213&gt; Conus characteristicus

&lt;400&gt; 65

Met Met Ser Lys Leu Gly Ala Leu Leu Thr Ile Cys Leu Leu Leu Phe  
1 5 10 15Ser Leu Thr Ala Val Pro Leu Asp Gly Asp Gln His Ala Asp Gln Pro  
20 25 30Ala Gln Arg Leu Gln Asp Arg Ile Pro Thr Glu Asp His Pro Leu Phe  
35 40 45Asp Pro Asn Lys Arg Cys Cys Pro Pro Val Ala Cys Asn Met Gly Cys  
50 55 60Lys Pro Cys Cys Gly  
65

&lt;210&gt; 66

&lt;211&gt; 15

&lt;212&gt; PRT

&lt;213&gt; Conus characteristicus

&lt;220&gt;

&lt;221&gt; PEPTIDE

&lt;222&gt; (1)..(15)

&lt;223&gt; Xaa at residue 3, 4 and 13 is Pro or Hyp

&lt;400&gt; 66

Cys Cys Xaa Xaa Val Ala Cys Asn Met Gly Cys Lys Xaa Cys Cys

20

1 5 10 15

&lt;210&gt; 67

&lt;211&gt; 293

&lt;212&gt; DNA

&lt;213&gt; Conus characteristicus

&lt;400&gt; 67

caagagggtt cgatagcagt tcatgatgtc taaaactggga gccttggta ccatctgtct 60  
 acttctgttt tccctaactg ctgttccgct ggatggagat caacatgcag accaacacctgc 120  
 agagcgtctg catgaccgc ttccaaactga aaatcatccc ttatatgatc ccgtcaaacg 180  
 gtgttgcgtt gattcggaaat gcgactattc ttgctggct tgctgtatgt ttggataacc 240  
 tttgttatacg cggcctcatc aagtgtctaa tgaataagta aaacgattgc agt 293

&lt;210&gt; 68

&lt;211&gt; 71

&lt;212&gt; PRT

&lt;213&gt; Conus characteristicus

&lt;400&gt; 68

Met	Met	Ser	Lys	Leu	Gly	Ala	Leu	Leu	Thr	Ile	Cys	Leu	Leu	Leu	Phe
1				5				10					15		

Ser	Leu	Thr	Ala	Val	Pro	Leu	Asp	Gly	Asp	Gln	His	Ala	Asp	Gln	Pro
			20			25					30				

Ala	Glu	Arg	Leu	His	Asp	Arg	Leu	Pro	Thr	Glu	Asn	His	Pro	Leu	Tyr
			35			40				45					

Asp	Pro	Val	Lys	Arg	Cys	Cys	Asp	Asp	Ser	Glu	Cys	Asp	Tyr	Ser	Cys
		50			55				60						

Trp	Pro	Cys	Cys	Met	Phe	Gly
65				70		

&lt;210&gt; 69

&lt;211&gt; 17

&lt;212&gt; PRT

&lt;213&gt; Conus characteristicus

&lt;220&gt;

&lt;221&gt; PEPTIDE

&lt;222&gt; (1)..(17)

<223> Xaa at residue 6 is Glu or gamma-carboxy Glu; Xaa at residue 13 is Pro or Hyp; Xaa at residue 12 is Trp or bromo-Trp; Xaa at residue 9 is Tyr, 125I-Tyr, mono-iodo-Tyr, di-iodo-Tyr, O-sulpho-Tyr or O-phospho-Ty

&lt;400&gt; 69

Cys	Cys	Asp	Asp	Ser	Xaa	Cys	Asp	Xaa	Ser	Cys	Xaa	Cys	Cys	Met
1					5				10			15		

Phe

&lt;210&gt; 70

&lt;211&gt; 232

&lt;212&gt; DNA

&lt;213&gt; Conus characteristicus

&lt;400&gt; 70

ggatccatga	tgtctaaact	gggagtcctt	ttgaccatct	gtctgcttct	gtttccctt	60
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actgctgttc	cgctggatgg	agatcaacct	gcagaccgac	ctgcagagcg	taagcaggac	120
gtttcatctg	aacagcatcc	cttctttgat	cccgtaaac	ggtgttgcgg	ccgggtttac	180
atggatgca	tcccttgg	cttttaacgt	gttcatgacc	aacttctcg	ag	232

<210> 71  
 <211> 68  
 <212> PRT  
 <213> Conus characteristicus

<400> 71  
 Gly Ser Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu  
 1 5 10 15

Leu Phe Pro Leu Thr Ala Val Pro Leu Asp Gly Asp Gln Pro Ala Asp  
 20 25 30

Arg Pro Ala Glu Arg Lys Gln Asp Val Ser Ser Glu Gln His Pro Phe  
 35 40 45

Phe Asp Pro Val Lys Arg Cys Cys Arg Arg Cys Tyr Met Gly Cys Ile  
 50 55 60

Pro Cys Cys Phe  
 65

<210> 72  
 <211> 14  
 <212> PRT  
 <213> Conus characteristicus

<220>  
 <221> PEPTIDE  
 <222> (1)...(14)  
 <223> Xaa at residue 11 is Pro or Hyp; Xaa at residue 6 is Tyr, 125I-Ty  
 r, mono-iodo-Tyr, di-iodo-Tyr, O-sulpho-Tyr or O-phospho-Ty

<400> 72  
 Cys Cys Arg Arg Cys Xaa Met Gly Cys Ile Xaa Cys Cys Phe  
 1 5 10

<210> 73  
 <211> 323  
 <212> DNA  
 <213> Conus circumcisus

<400> 73  
 caagaaggat cgatagcagt tcatgatgtc taaaactgggg gtattgttga ccatctgtct 60  
 gcttctgttt ccccttactg ctcttccaaat ggtggagat caacctgcag accaacctgc 120  
 agatcgatg caggacgaca tttcatctga gcagtatccc ttgtttgata agagacgaaa 180  
 gtgttgcggc aaagacgggc catgccccaa atatttcaaa gacaattta ttgtggttg 240  
 ttgttaatg acaacgtgtc gatgaccaac ttctgttatca cgattcgcca agtgcattaa 300  
 tgaataagta aaatgattgc agt 323

<210> 74  
 <211> 74  
 <212> PRT  
 <213> Conus circumcisus

<400> 74  
 Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15

Pro Leu Thr Ala Leu Pro Met Asp Gly Asp Gln Pro Ala Asp Gln Pro  
 20 25 30

Ala Asp Arg Met Gln Asp Asp Ile Ser Ser Glu Gln Tyr Pro Leu Phe  
 35 40 45

Asp Lys Arg Arg Lys Cys Cys Gly Lys Asp Gly Pro Cys Pro Lys Tyr  
 50 55 60

Phe Lys Asp Asn Phe Ile Cys Gly Cys Cys  
 65 70

<210> 75

<211> 23

<212> PRT

<213> Conus circumcisus

<220>

<221> PEPTIDE

<222> (1)..(23)

<223> Xaa at residue 9 and 11 is Pro or Hyp; Xaa at residue 13 is Tyr,  
 125I-Tyr, mono-iodo-Tyr, di-iodo-Tyr, O-sulpho-Tyr or O-phospho-T  
 y

<400> 75

Arg Lys Cys Cys Gly Lys Asp Gly Xaa Cys Xaa Lys Xaa Phe Lys Asp  
 1 5 10 15

Asn Phe Ile Cys Gly Cys Cys  
 20

<210> 76

<211> 293

<212> DNA

<213> Conus dalli

<400> 76

caagaggat cgatagcagt tcatgatgtc taaaactggga gccttgttga ccatctgtct 60

acttctgttt tccctaactg ctgttccgct ggatggagat caacatgcag accaacctgc 120

agagcgtctg caggaccgcc ttccaactga aaatcatccc ttatatgatc ccgtcaaacg 180

gtgttgcgat gattcggaat gcgactattc ttgctggct tgctgtattt tatcataacc 240

tttggtatcg cggcctcatc aagtgtcaaa tgaataagta aaatgattgc agt 293

<210> 77

<211> 71

<212> PRT

<213> Conus dalli

<400> 77

Met Met Ser Lys Leu Gly Ala Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15

Ser Leu Thr Ala Val Pro Leu Asp Gly Asp Gln His Ala Asp Gln Pro  
 20 25 30

Ala Glu Arg Leu Gln Asp Arg Leu Pro Thr Glu Asn His Pro Leu Tyr  
 35 40 45

Asp Pro Val Lys Arg Cys Cys Asp Asp Ser Glu Cys Asp Tyr Ser Cys  
 50 55 60

Trp Pro Cys Cys Ile Leu Ser  
 65 70

<210> 78  
 <211> 18  
 <212> PRT  
 <213> Conus dalli

<220>  
 <221> PEPTIDE  
 <222> (1)...(18)  
 <223> Xaa at residue 6 is Glu or gamma-carboxy Glu; Xaa at residue 13 is Pro or Hyp; Xaa at residue 12 is Trp or bromo-Trp; Xaa at residue 9 is Tyr, 125I-Tyr, mono-iodo-Tyr, di-iodo-Tyr, O-sulpho-Tyr or O-phospho-Tyr

<400> 78  
 Cys Cys Asp Asp Ser Xaa Cys Asp Xaa Ser Cys Xaa Xaa Cys Cys Ile  
 1 5 10 15

Leu Ser

<210> 79  
 <211> 299  
 <212> DNA  
 <213> Conus dalli

<400> 79  
 caagaggggat cgatagcagt tcatgatgtc taaaactggga gtcttggta ccatttgtct 60  
 acttctgttt ccccttactg ctgttccact ggatggagat cagcctgcag accgacctgc 120  
 agagcgtatg caggacggca tttcatctga acatcatcca tttttgatt ccgtcaaaaa 180  
 gaaacaacag tggcccgc cggtggcatg caacatggga tgcgagcctt gttgtggatg 240  
 accagctttg ttatcgccgc tcatgaagtgc tcctaatgaa taagtaaaac gattgcagt 299

<210> 80  
 <211> 72  
 <212> PRT  
 <213> Conus dalli

<400> 80  
 Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15

Pro Leu Thr Ala Val Pro Leu Asp Gly Asp Gln Pro Ala Asp Arg Pro  
 20 25 30

Ala Glu Arg Met Gln Asp Gly Ile Ser Ser Glu His His Pro Phe Phe  
 35 40 45

Asp Ser Val Lys Lys Gln Gln Cys Cys Pro Pro Val Ala Cys Asn  
 50 55 60

Met Gly Cys Glu Pro Cys Cys Gly  
 65 70

<210> 81  
 <211> 17

<212> PRT  
 <213> Conus dalli  
  
 <220>  
 <221> PEPTIDE  
 <222> (1)...(17)  
 <223> Xaa at residue 1 is Gln or pyro-Glu; Xaa at residue 14 is Glu or  
 gamma-carboxy Glu; Xaa at residue 5, 6 and 15 is Pro or Hyp  
  
 <400> 81  
 Xaa Gln Cys Cys Xaa Xaa Val Ala Cys Asn Met Gly Cys Xaa Xaa Cys  
 1 5 10 15  
  
 Cys  
  
 <210> 82  
 <211> 290  
 <212> DNA  
 <213> Conus dalli  
  
 <400> 82  
 caagaaggat cgatagcagt tcatgatgtc taaaactggga gtcttggta tcataatgtct 60  
 atttctgttt ccccttactg ctgttcagct caatggagat cagcctgcag accaatctgc 120  
 agagcgtatg caggacaaaa tttcatctga acatcatccc tttttgtatc ccgtcaaaacg 180  
 ttgttgcac ac gccccctttt gcccgttcgg atgcacgcct tttttttttt gaccagcttt 240  
 gttatcgcgg cctcatcaag tgtctaatga ataagtaaaa tgattgcagt 290  
 <210> 83  
 <211> 69  
 <212> PRT  
 <213> Conus dalli  
  
 <400> 83  
 Met Met Ser Lys Leu Gly Val Leu Leu Ile Ile Cys Leu Phe Leu Phe  
 1 5 10 15  
  
 Pro Leu Thr Ala Val Gln Leu Asn Gly Asp Gln Pro Ala Asp Gln Ser  
 20 25 30  
  
 Ala Glu Arg Met Gln Asp Lys Ile Ser Ser Glu His His Pro Phe Phe  
 35 40 45  
  
 Asp Pro Val Lys Arg Cys Cys Asn Ala Gly Phe Cys Arg Phe Gly Cys  
 50 55 60  
  
 Thr Pro Cys Cys Trp  
 65  
  
 <210> 84  
 <211> 16  
 <212> PRT  
 <213> Conus dalli  
  
 <220>  
 <221> PEPTIDE  
 <222> (1)...(16)  
 <223> Xaa at residue 13 is Pro or Hyp; Xaa at residue 16 is Trp or brom  
 o-Tr  
  
 <400> 84  
 Cys Cys Asn Ala Gly Phe Cys Arg Phe Gly Cys Thr Xaa Cys Cys Xaa  
 1 5 10 15

<210> 85  
 <211> 288  
 <212> DNA  
 <213> *Conus distans*

<400> 85  
 caagaggat cgatagcagt tcatgatgtc taaaactggga gtcttgctga ccatcttct 60  
 gcttctgtt ccccttactg ctgttccgct gcatggagat caaccgcag acggacttgc 120  
 agagcgcatg caggacgaca gttcagctgc actgattaga gactggcttc ttcaaaccgg 180  
 acagtgttgt gtgcacatccat gcccacatgcac gccttgcgtt agatgaccag ctttgcac 240  
 gcccgcgtt acgttatcta atgaaataagt aagtaaaacg attgcagt 288

<210> 86  
 <211> 67  
 <212> PRT  
 <213> *Conus distans*

<400> 86  
 Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Phe Leu Leu Leu Phe  
 1 5 10 15

Pro Leu Thr Ala Val Pro Leu Asp Gly Asp Gln Pro Ala Asp Gly Leu  
 20 25 30

Ala Glu Arg Met Gln Asp Asp Ser Ser Ala Ala Leu Ile Arg Asp Trp  
 35 40 45

Leu Leu Gln Thr Arg Gln Cys Cys Val His Pro Cys Pro Cys Thr Pro  
 50 55 60

Cys Cys Arg  
 65

<210> 87  
 <211> 14  
 <212> PRT  
 <213> *Conus distans*

<220>  
 <221> PEPTIDE  
 <222> (1)...(14)  
 <223> Xaa at residue 1 is Gln or pyro-Glu; Xaa at residue 6, 8 and 11 is Pro or Hy

<400> 87  
 Xaa Cys Cys Val His Xaa Cys Xaa Cys Thr Xaa Cys Cys Arg  
 1 5 10

<210> 88  
 <211> 303  
 <212> DNA  
 <213> *Conus ermineus*

<400> 88  
 acctcaagag ggatcgatcg cagttcatga tgtctaaact gggagccttg ttgaccatct 60  
 gtctgcatttct gtttccattt actgctttc tggatggatgg agatcagcct gcagaccgac 120  
 ctgcagagcg tacggaggat gacatttcat ctgactacat tccctgttgc agttggccat 180

gcccccgata ctccaacggt aaacttgtt gttttgttgccttggatga taatgtgtt 240  
 atgaccaact ttgttatcac ggctacgtca agtgtctact gaataagtaa aatgattgca 300  
 gta 303  
 <210> 89  
 <211> 67  
 <212> PRT  
 <213> *Conus ermineus*  
 <400> 89  
 Met Met Ser Lys Leu Gly Ala Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15  
 Pro Ile Thr Ala Leu Leu Met Asp Gly Asp Gln Pro Ala Asp Arg Pro  
 20 25 30  
 Ala Glu Arg Thr Glu Asp Asp Ile Ser Ser Asp Tyr Ile Pro Cys Cys  
 35 40 45  
 Ser Trp Pro Cys Pro Arg Tyr Ser Asn Gly Lys Leu Val Cys Phe Cys  
 50 55 60  
 Cys Leu Gly  
 65  
 <210> 90  
 <211> 20  
 <212> PRT  
 <213> *Conus ermineus*  
 <220>  
 <221> PEPTIDE  
 <222> (1)..(20)  
 <223> Xaa at residue 5 and 7 is Pro or Hyp; Xaa at residue 4 is Trp or  
 bromo-Trp; Xaa at residue 9 is Tyr, 125I-Tyr, mono-iodo-Tyr, di-  
 iodo-Tyr, O-sulpho-Tyr or O-phospho-Ty  
 <400> 90  
 Cys Cys Ser Xaa Xaa Cys Xaa Arg Xaa Ser Asn Gly Lys Leu Val Cys  
 1 5 10 15  
 Phe Cys Cys Leu  
 20  
 <210> 91  
 <211> 241  
 <212> DNA  
 <213> *Conus generalis*  
 <400> 91  
 ggatccatga tgtctaaact gggagtcttg ttgaccatct gtctggttct gtttcccctt 60  
 actgctcttc cactggatgg agaacaacct gtagaccgac atgccgagca tatgcaggat 120  
 gacaattcag ctgcacagaa cccctgggtt attgccatca gacagtgttgcacgttctgc 180  
 aactttggat gccaaccttg ttgcctcacc tgataacgtg ttgatgacca aacttctcga 240  
 g 241  
 <210> 92  
 <211> 70  
 <212> PRT

&lt;213&gt; Conus generalis

<400> 92  
 Gly Ser Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Val  
 1 5 10 15  
 Leu Phe Pro Leu Thr Ala Leu Pro Leu Asp Gly Glu Gln Pro Val Asp  
 20 25 30  
 Arg His Ala Glu His Met Gln Asp Asp Asn Ser Ala Ala Gln Asn Pro  
 35 40 45  
 Trp Val Ile Ala Ile Arg Gln Cys Cys Thr Phe Cys Asn Phe Gly Cys  
 50 55 60  
 Gln Pro Cys Cys Leu Thr  
 65 70

<210> 93  
<211> 16  
<212> PRT  
<213> Conus generalis

<220>  
<221> PEPTIDE  
<222> (1)...(16)  
<223> Xaa at residue 1 is Gln or pyro-Glu; Xaa at residue 12 is Pro or  
 Hy

<400> 93  
 Xaa Cys Cys Thr Phe Cys Asn Phe Gly Cys Gln Xaa Cys Cys Cys Leu Thr  
 1 5 10 15

<210> 94  
<211> 241  
<212> DNA  
<213> Conus generalis

<400> 94  
 ggcatccatga tgcctaaact gggagtcttg ttgaccatct gtctggttct gtttccctt 60  
 actgctcttc cactggatgg agaacaacccat gtagaccgac atgcccggca tatgcaggat 120  
 gacaattcag ctgcacagaa cccctgggtt attgccatca gacagtgttg cacgttctgc 180  
 aactttggat gccagccttg ttgcgtcccc tgataacgtg ttgatgacca actttctcga 240  
 g 241

<210> 95  
<211> 70  
<212> PRT  
<213> Conus generalis

<400> 95  
 Gly Ser Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Val  
 1 5 10 15

Leu Phe Pro Leu Thr Ala Leu Pro Leu Asp Gly Glu Gln Pro Val Asp  
 20 25 30

Arg His Ala Glu His Met Gln Asp Asp Asn Ser Ala Ala Gln Asn Pro  
 35 40 45

Trp Val Ile Ala Ile Arg Gln Cys Cys Thr Phe Cys Asn Phe Gly Cys

50

55

60

Gln Pro Cys Cys Val Pro  
 65 70

&lt;210&gt; 96

&lt;211&gt; 16

&lt;212&gt; PRT

&lt;213&gt; Conus generalis

&lt;220&gt;

&lt;221&gt; PEPTIDE

&lt;222&gt; (1)..(16)

&lt;223&gt; Xaa at residue 1 is Gln or pyro-Glu; Xaa at residue 12 and 16 is Pro or Hy

&lt;400&gt; 96

Xaa Cys Cys Thr Phe Cys Asn Phe Gly Cys Gln Xaa Cys Cys Cys Val Xaa  
 1 5 10 15

&lt;210&gt; 97

&lt;211&gt; 862

&lt;212&gt; DNA

&lt;213&gt; Conus geographus

&lt;400&gt; 97

gtcgactcta gaggatccga caacaaagag tcaaccccac tgccacgtca agagcgaagc 60  
 gccacagcta agacaagagg gatcgatagc agttcatgtat gtctaaactg ggagtcttgt 120  
 tgaccatctg tctgcttctg tttccctta ctgctttcc gatggatgga gatgaacctg 180  
 caaaccgacc tgtcgagcgt atgcaggaca acatttcatc tgagcagttat cccttgttg 240  
 agaagagacg agattgttgc actccgcccga agaaatgcaa agaccgacaa tgcaaacc 300  
 agagatgttgc cgctggacga taacgttttg atgaccaact ttatcacggc tacgtcaagt 360  
 gtttagtcaa taagtaaaat gattgcagtc ttgctcagat ttgctttgt gtttggtct 420  
 aaagatcaat gaccaaaccg ttgttttgat gcggattgtc atatatttct cgattccat 480  
 ccaacactag atgatTTat cacgatagat taatTTcta tcaatgcctt gatTTTcgt 540  
 ctgtcatatc agtttggat atatTTattt ttctgtcaact gtctacacaa acgcatgcat 600  
 gcacgcacatgc acgcacacac gcacgcacgc tcgcacaaac atgcgcgc acgcacacac 660  
 acacacacac acacaaacac acacacaacg aatcacacaa ttattgacat tattttat 720  
 ttcatgtat tattgttat tcgtttgtt gtttttagaa tagtttggagg ccgttttttt 780  
 ggattttatggat gaaactgcttt attgtataacg agtacttcgt gctttgaaac actgctgaaa 840  
 ataaaaacaaaa cactgacgta gc 862

&lt;210&gt; 98

&lt;211&gt; 75

&lt;212&gt; PRT

&lt;213&gt; Conus geographus

&lt;400&gt; 98

Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15

Pro Leu Thr Ala Leu Pro Met Asp Gly Asp Glu Pro Ala Asn Arg Pro  
 20 25 30

Val Glu Arg Met Gln Asp Asn Ile Ser Ser Glu Gln Tyr Pro Leu Phe  
 35 40 45

Glu Lys Arg Arg Asp Cys Cys Thr Pro Pro Lys Lys Cys Lys Asp Arg  
 50 55 60

Gln Cys Lys Pro Gln Arg Cys Cys Ala Gly Arg  
 65 70 75

<210> 99

<211> 22

<212> PRT

<213> Conus geographus

<220>

<221> PEPTIDE

<222> (1)..(22)

<223> Xaa at residue 6, 7 and 17 is Pro or Hyp

<400> 99

Arg Asp Cys Cys Thr Xaa Xaa Lys Lys Cys Lys Asp Arg Gln Cys Lys  
 1 5 10 15

Xaa Gln Arg Cys Cys Ala  
 20

<210> 100

<211> 860

<212> DNA

<213> Conus geographus

<400> 100

ggccagacga caacaaagag tcaacccac tgccacgtca agagcgaagc gccacagcta 60

agacaagagg gatcgatgc agttcatgat gtctaaactg ggagtcttgt tgaccatctg 120

tctgcttctg tttccctta ctgcttcc gatggatgga gatgaacctg caaaccgacc 180

tgtcgacgt atgcaggaca acatttcatc tgagcgtat cccttgtttg agaagagacg 240

agattttgc actccgcccga ggaaatgcaa agaccgacga tgcaaaccga tgaaatgtt 300

cgctggacga taaegtgtt atgaccaact ttatcaeggc tagtcagtg ttttgtaat 360

aagtaaaatg attgcagtct tgctcagatt gctttgtgt tttggctaa gatcaatgac 420

caaaccgtt ttttgcgtcg gattgtcata tatttcgtca ttccaaatcca acactagatg 480

atttaatcac gatagattaa ttttctatca atgcctgtat ttttcgtctg tcataatcgt 540

tttgtttata tttatgtt cgtcaactgtc tacacaaacg catgcgtgc cgcgtgcacg 600

cacacacgca cgcacgctcg cacaacatg cgcgcgcacg cacacacaca cacacacaca 660

aacacacaca cgaagcaatc acacaattag ttgacattat ttatatttc attgtatgtat 720

ttgttattcg ttgcgtgtt tttagaatag tttgaggccg tcttttgaa ttatgttggaa 780

ctgcatttatt gtatacgagt acttcgtgtt ttgaaacact gctgaaaata aaacaaacac 840

tgacgttagca aaaaaaaaaaa 860

30

<210> 101  
 <211> 75  
 <212> PRT  
 <213> Conus geographus

<400> 101  
 Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15

Pro Leu Thr Ala Leu Pro Met Asp Gly Asp Glu Pro Ala Asn Arg Pro  
 20 25 30

Val Glu Arg Met Gln Asp Asn Ile Ser Ser Glu Gln Tyr Pro Leu Phe  
 35 40 45

Glu Lys Arg Arg Asp Cys Cys Thr Pro Pro Arg Lys Cys Lys Asp Arg  
 50 55 60

Arg Cys Lys Pro Met Lys Cys Cys Ala Gly Arg  
 65 70 75

<210> 102  
 <211> 22  
 <212> PRT  
 <213> Conus geographus

<220>  
 <221> PEPTIDE  
 <222> (1)..(22)  
 <223> Xaa at residue 6, 7 and 17 is Pro or Hyp

<400> 102  
 Arg Asp Cys Cys Thr Xaa Xaa Arg Lys Cys Lys Asp Arg Arg Cys Lys  
 1 5 10 15

Xaa Met Lys Cys Cys Ala  
 20

<210> 103  
 <211> 22  
 <212> PRT  
 <213> Conus geographus

<220>  
 <221> PEPTIDE  
 <222> (1)..(22)  
 <223> Xaa at residue 6, 7 and 17 is Pro or Hyp

<400> 103  
 Arg Asp Cys Cys Thr Xaa Xaa Lys Lys Cys Lys Asp Arg Arg Cys Lys  
 1 5 10 15

Xaa Leu Lys Cys Cys Ala  
 20

<210> 104  
 <211> 321  
 <212> DNA  
 <213> Conus gloriamaris

<400> 104  
 ctcactatacg aattcgagc tcggcacacg ggatcgatag cagttcatga tgtctaaact 60  
 gggagccttg ttgaccatct gtctacttct gttttcccta actgctgttc cgctggatgg 120

agatcaacat gcagaccaac ctgcagagcg tctgcacatgac cgccttccaa ctgaaaatca 180  
 tcccttataat gatcccgatca aacgggtttg cgatgattcg gaatgcgact attcttgctg 240  
 gccttgcgtgt atgtttggat aacccttggat atcgcggcct cgataagtgt ctaatgaata 300  
 agtaaaaacga ttgcagtagg c 321  
  
 <210> 105  
 <211> 71  
 <212> PRT  
 <213> Conus gloriamaris  
  
 <400> 105  
 Met Met Ser Lys Leu Gly Ala Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15  
  
 Ser Leu Thr Ala Val Pro Leu Asp Gly Asp Gln His Ala Asp Gln Pro  
 20 25 30  
  
 Ala Glu Arg Leu His Asp Arg Leu Pro Thr Glu Asn His Pro Leu Tyr  
 35 40 45  
  
 Asp Pro Val Lys Arg Cys Cys Asp Asp Ser Glu Cys Asp Tyr Ser Cys  
 50 55 60  
  
 Trp Pro Cys Cys Met Phe Gly  
 65 70  
  
 <210> 106  
 <211> 17  
 <212> PRT  
 <213> Conus gloriamaris  
  
 <220>  
 <221> PEPTIDE  
 <222> (1)..(17)  
 <223> Xaa at residue is 6 Glu or gamma-carboxy Glu; Xaa at residue 13 is  
 Pro or Hyp; Xaa at residue 12 is Trp or bromo-Trp; Xaa at residue 9 is  
 Tyr, 125I-Tyr, mono-iodo-Tyr, di-iodo-Tyr, O-sulpho-Tyr or  
 O-phospho-Ty  
  
 <400> 106  
 Cys Cys Asp Asp Ser Xaa Cys Asp Xaa Ser Cys Xaa Xaa Cys Cys Met  
 1 5 10 15  
  
 Phe  
  
 <210> 107  
 <211> 257  
 <212> DNA  
 <213> Conus gloriamaris  
  
 <400> 107  
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 tgctgttccg ctggatggag atcaacactgc agaccgatat gcagagcgta tgcaggacga 120  
 catttcatctt gaacatcatc ccatgtttga tgccgtcaga ggggtttgcc atctgttggc 180  
 atgccgccttc ggatgctcgc cttgttggat gtgatcagct ttgttatcgc ggcctcatca 240  
  
 agtgactcta atgcaaa 257  
 <210> 108  
 <211> 69

<212> PRT  
 <213> Conus gloriamaris

<400> 108  
 Met Met Ser Lys Leu Gly Val Leu Leu Ile Ile Cys Leu Leu Leu Phe  
 1 5 10 15

Pro Leu Thr Ala Val Pro Leu Asp Gly Asp Gln Pro Ala Asp Arg Tyr  
 20 25 30

Ala Glu Arg Met Gln Asp Asp Ile Ser Ser Glu His His Pro Met Phe  
 35 40 45

Asp Ala Val Arg Gly Cys Cys His Leu Leu Ala Cys Arg Phe Gly Cys  
 50 55 60

Ser Pro Cys Cys Trp  
 65

<210> 109

<211> 17

<212> PRT

<213> Conus gloriamaris

<220>

<221> PEPTIDE

<222> (1)..(17)

<223> Xaa at residue 14 is Pro or Hyp; Xaa at residue 17 is Trp or brom  
 o-Tr

<400> 109

Gly Cys Cys His Leu Leu Ala Cys Arg Phe Gly Cys Ser Xaa Cys Cys  
 1 5 10 15

Xaa

<210> 110

<211> 471

<212> DNA

<213> Conus gloriamaris

<400> 110

gagacgacaa ggaacagtca accccacagc cacgccaaga gcagacagcc acagctacgt 60

gaagaagggt ggagagaggt tcgtgatgtt gaaaatggga gtggtgctat tcatttcct 120

ggtaactgtt cccctggcaa cgctccagct ggatgcagat caacctgttag aacgatatgc 180

ggagaacaaa cagctcctca acccagatga aaggaggaa atcatattgc atgctctggg 240

gacgcgatgc ttttcttggg atgtgtgcga ccacccaggt tgtacttgct gggcggtta 300

gccccgaaaca tccatggcgc tgtgtgggc gttttatcc aacaacgaca gcgtttgttg 360

atttcatgtt tcaattgcgcc cacgtctctt gtctaagaat gacgaacatg attgcactct 420

ggttcagatt tcgtgttctt ttctgacaat aaatgacaaa actccaaaaa a 471

<210> 111

<211> 71

<212> PRT

<213> Conus gloriamaris

<400> 111

Met Leu Lys Met Gly Val Val Leu Phe Ile Phe Leu Val Leu Phe Pro

33

1	5	10	15												
Leu	Ala	Thr	Leu	Gln	Leu	Asp	Ala	Asp	Gln	Pro	Val	Glu	Arg	Tyr	Ala
			20			25				30					

Glu	Asn	Lys	Gln	Leu	Leu	Asn	Pro	Asp	Glu	Arg	Arg	Glu	Ile	Ile	Leu
			35			40			45						

His	Ala	Leu	Gly	Thr	Arg	Cys	Cys	Ser	Trp	Asp	Val	Cys	Asp	His	Pro
			50			55			60						

Ser	Cys	Thr	Cys	Cys	Gly	Gly
65		50		70		

&lt;210&gt; 112

&lt;211&gt; 16

&lt;212&gt; PRT

&lt;213&gt; Conus gloriamaris

&lt;220&gt;

&lt;221&gt; PEPTIDE

&lt;222&gt; (1)..(16)

&lt;223&gt; Xaa at residue 10 is Pro or Hyp; Xaa at residue 4 is Trp or bromo-Tr

&lt;400&gt; 112

Cys	Cys	Ser	Xaa	Asp	Val	Cys	Asp	His	Xaa	Ser	Cys	Thr	Cys	Cys	Gly
1			5			10			15						

&lt;210&gt; 113

&lt;211&gt; 304

&lt;212&gt; DNA

&lt;213&gt; Conus laterculatus

&lt;400&gt; 113

cgacctcaag aaggatcgat agcagttcat gatgtctaaa ctgggagtct tggaccat 60

ctgtctgctt ctgtttcccc ttactgctct tccgatggat ggagatcaac ctgcagaccg 120

acctgcagag cgtatgcagg acgtttcatc tgaacagcat ccctgtatg atcccgtaa 180

acgggtttgc gactggccat gcagcgatg catccctgt tgctaatagt aacaacgtgt 240

tgataaccaa ctttcttacc acgactacgt caagtgtcta atgaataagt aaaatgattg 300

cagt 304

&lt;210&gt; 114

&lt;211&gt; 65

&lt;212&gt; PRT

&lt;213&gt; Conus laterculatus

&lt;400&gt; 114

Met	Met	Ser	Lys	Leu	Gly	Val	Leu	Leu	Thr	Ile	Cys	Leu	Leu	Phe
1				5			10		15					

Pro	Leu	Thr	Ala	Leu	Pro	Met	Asp	Gly	Asp	Asp	Gln	Pro	Ala	Asp	Arg	Pro
			20			25		30								

Ala	Glu	Arg	Met	Gln	Asp	Val	Ser	Ser	Glu	Gln	His	Pro	Leu	Tyr	Asp
			35			40			45						

Pro	Val	Lys	Arg	Cys	Cys	Asp	Trp	Pro	Cys	Ser	Gly	Cys	Ile	Pro	Cys
			50			55			60						

Cys

65

<210> 115  
 <211> 13  
 <212> PRT  
 <213> Conus laterculatus

<220>  
 <221> PEPTIDE  
 <222> (1)..(13)  
 <223> Xaa at residue 5 and 11 is Pro or Hyp; Xaa at residue 4 is Trp or  
 bromo-Trp

<400> 115  
 Cys Cys Asp Xaa Xaa Cys Ser Gly Cys Ile Xaa Cys Cys  
 1 5 10

<210> 116  
 <211> 313  
 <212> DNA  
 <213> Conus laterculatus

<400> 116  
 cgacacctaa aaggatcgat agcagttcat gatgtctaaa ctgggagtct tggaccat 60  
 ctgtctgctt ctgtttcccc ttactgtctt ggtggagat caacctgcag accgacttgc 120  
 agagcgatag caggacgaca ttcatctga gcagcatccc ttgaaaaga gacgagactg 180  
 ttgcacacct ccgaagaaat gcagagaccg acaatgcaaa cctgcacggtt gttgcggagg 240  
 ataacgtgtt gatgaccaac tttgttatca cggctacgtc aagtgtctag tgaataagta 300  
 aaacgattgc agt 313

<210> 117  
 <211> 71  
 <212> PRT  
 <213> Conus laterculatus

<400> 117  
 Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15

Pro Leu Thr Ala Leu Asp Gly Asp Gln Pro Ala Asp Arg Leu Ala Glu  
 20 25 30

Arg Met Gln Asp Asp Ile Ser Ser Glu Gln His Pro Phe Glu Lys Arg  
 35 40 45

Arg Asp Cys Cys Thr Pro Pro Lys Lys Cys Arg Asp Arg Gln Cys Lys  
 50 55 60

Pro Ala Arg Cys Cys Gly Gly  
 65 70

<210> 118  
 <211> 22  
 <212> PRT  
 <213> Conus laterculatus

<220>  
 <221> PEPTIDE  
 <222> (1)..(22)  
 <223> Xaa at residue 6, 17 and 17 is Pro or Hyp

<400> 118  
 Arg Asp Cys Cys Thr Xaa Xaa Lys Lys Cys Arg Asp Arg Gln Cys Lys  
 1 5 10 15

Xaa Ala Arg Cys Cys Gly  
 20

<210> 119  
 <211> 314  
 <212> DNA  
 <213> Conus laterculatus

<400> 119  
 gggatcgata gcagttcatg atgtctaaac tgggagtctt gttgaccatc tgtctgcttc 60  
 tttttccct tactgtcttt ccgatggatg gagatcaact tgcacgccga tctgcagagc 120  
 gtagcagga caacatttca tctgagcagc atcaccttt tgaaaagaga cgaccaccat 180  
 gttgcaccta tgacggagt tgcctaaaag aatcatgcat gcttaaagct ttttgcggat 240  
 gataacgtgt tgatgaccaa ctttgttatac acggctactc aagtgtctaa tgaataagta 300  
 aaatgattgc agta 314

<210> 120  
 <211> 74  
 <212> PRT  
 <213> Conus laterculatus

<400> 120  
 Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15

Pro Leu Thr Ala Leu Pro Met Asp Gly Asp Gln Leu Ala Arg Arg Ser  
 20 25 30

Ala Glu Arg Met Gln Asp Asn Ile Ser Ser Glu Gln His His Leu Phe  
 35 40 45

Glu Lys Arg Arg Pro Pro Cys Cys Thr Tyr Asp Gly Ser Cys Leu Lys  
 50 55 60

Glu Ser Cys Met Arg Lys Ala Cys Cys Gly  
 65 70

<210> 121  
 <211> 22  
 <212> PRT  
 <213> Conus laterculatus

<220>  
 <221> PEPTIDE  
 <222> (1)..(22)  
 <223> Xaa at residue 14 is Glu or gamma-carboxy Glu; Xaa at residue 2 and 3 is Pro or Hyp; Xaa at residue 7 is Tyr, 125I-Tyr, mono-iodo-Tyr, di-iodo-Tyr, O-sulpho-Tyr or O-phospho-Ty

<400> 121  
 Arg Xaa Xaa Cys Cys Thr Xaa Asp Gly Ser Cys Leu Lys Xaa Ser Cys  
 1 5 10 15

Met Arg Lys Ala Cys Cys  
 20

<210> 122  
 <211> 314  
 <212> DNA  
 <213> *Conus laterculatus*

<400> 122  
 gggatcgata gcagttcatg atgtctaaac tgggagtctt gttgaccacc tgtctgtttc 60  
 tgtttccct tactgctctt ccgatggatg gagatcaact tgcacgccga cctgcagagc 120  
 gtatgcagga caacatttca tctgagcagc atcccttctt tgaaaggaga cgaccaccat 180  
 gttgcaccta tgacgggagt tgcctaaaag aatcatgcaa gcttaagct tggat 240  
 aataacgtgt tggatgaccaa ctttggatc acggctactc aagtgtctaa tgaataagta 300  
 aatgattgc agta 314

<210> 123  
 <211> 74  
 <212> PRT  
 <213> *Conus laterculatus*

<400> 123  
 Met Met Ser Lys Leu Gly Val Leu Leu Thr Thr Cys Leu Leu Leu Phe  
 1 5 10 15

Pro Leu Thr Ala Leu Pro Met Asp Gly Asp Gln Leu Ala Arg Arg Pro  
 20 25 30

Ala Glu Arg Met Gln Asp Asn Ile Ser Ser Glu Gln His Pro Phe Phe  
 35 40 45

Glu Arg Arg Arg Pro Pro Cys Cys Thr Tyr Asp Gly Ser Cys Leu Lys  
 50 55 60

Glu Ser Cys Lys Arg Lys Ala Cys Cys Gly  
 65 70

<210> 124  
 <211> 22  
 <212> PRT  
 <213> *Conus laterculatus*

<220>  
 <221> PEPTIDE  
 <222> (1)..(22)  
 <223> Xaa at residue 14 is Glu or gamma-carboxy Glu; Xaa at residue 2 and 3 is Pro or Hyp; Xaa at residue 7 is Tyr, 125I-Tyr, mono-iodo-Tyr, di-iodo-Tyr, O-sulpho-Tyr or O-phospho-Ty

<400> 124  
 Arg Xaa Xaa Cys Cys Thr Xaa Asp Gly Ser Cys Leu Lys Xaa Ser Cys  
 1 5 10 15

Lys Arg Lys Ala Cys Cys  
 20

<210> 125  
 <211> 247  
 <212> DNA  
 <213> *Conus leopardus*

<400> 125

ggatccatga tgtctaaact gggagtcttg ttgaccgtct gtctgcttct gttcccccctt 60  
 actgctcttc ggctgggttgg agatcaacct gcagagcgac ctgcaaagcg tacgcaggac 120  
 gacattccag atggacagca tccgttaat gataggcaga taaactgttg cccgtggcca 180  
 tgccctagta catgccgcca tcaatgctgc cattaatgat aacgtgttga tgaccaactt 240  
 tctcgag 247

<210> 126  
 <211> 71  
 <212> PRT  
 <213> Conus leopardus

<400> 126  
 Gly Ser Met Met Ser Lys Leu Gly Val Leu Leu Thr Val Cys Leu Leu  
 1 5 10 15

Leu Phe Pro Leu Thr Ala Leu Arg Leu Val Gly Asp Gln Pro Ala Glu  
 20 25 30

Arg Pro Ala Lys Arg Thr Gln Asp Asp Ile Pro Asp Gly Gln His Pro  
 35 40 45

Leu Asn Asp Arg Gln Ile Asn Cys Cys Pro Trp Pro Cys Pro Ser Thr  
 50 55 60

Cys Arg His Gln Cys Cys His  
 65 70

<210> 127  
 <211> 19  
 <212> PRT  
 <213> Conus leopardus

<220>  
 <221> PEPTIDE  
 <222> (1)..(19)  
 <223> Xaa at residue 1 is Gln or pyro-Glu; Xaa at residue 6, 8 and 10 is Pro or Hyp; Xaa at residue 7 is Trp or bromo-Tr

<400> 127  
 Xaa Ile Asn Cys Cys Xaa Xaa Xaa Cys Xaa Ser Thr Cys Arg His Gln  
 1 5 10 15

Cys Cys His

<210> 128  
 <211> 244  
 <212> DNA  
 <213> Conus lividus

<400> 128  
 ggatccatga tgtctaaact gggagtcttg ttgaccgtct gtctgcttct gttcccccctt 60  
 actgctcttc ggctgggttag agatcaacct gcagagcgac ctgcaaagcg tacgcaggac 120  
 gacattccaa atggacagga tccgttaatt gataggcaga taaattgttg cccttggcca 180  
 tgccctgatt catgccacta tcaatgctgc cactgataac gtgttgatga ccaactttct 240  
 cgag 244

<210> 129

<211> 71  
 <212> PRT  
 <213> Conus lividus

<400> 129  
 Gly Ser Met Met Ser Lys Leu Gly Val Leu Leu Thr Val Val Cys Leu Leu  
 1 5 10 15

Leu Phe Pro Leu Thr Ala Leu Arg Leu Val Arg Asp Gln Pro Ala Glu  
 20 25 30

Arg Pro Ala Lys Arg Thr Gln Asp Asp Ile Pro Asn Gly Gln Asp Pro  
 35 40 45

Leu Ile Asp Arg Gln Ile Asn Cys Cys Pro Trp Pro Cys Pro Asp Ser  
 50 55 60

Cys His Tyr Gln Cys Cys His  
 65 70

<210> 130  
 <211> 19  
 <212> PRT  
 <213> Conus lividus

<220>  
 <221> PEPTIDE  
 <222> (1)..(19)  
 <223> Xaa at residue 1 is Gln or pyro-Glu; Xaa at residue 6, 8 and 10 is  
 Pro or Hyp; Xaa at residue 7 is Trp or bromo-Trp; Xaa at residue 15 is  
 Tyr, 125I-Tyr, mono-iodo-Tyr, di-iodo-Tyr, O-sulpho-Tyr or  
 O-phospho-Ty

<400> 130  
 Xaa Ile Asn Cys Cys Xaa Xaa Xaa Cys Xaa Asp Ser Cys His Xaa Gln  
 1 5 10 15

Cys Cys His

<210> 131  
 <211> 275  
 <212> DNA  
 <213> Conus lynceus

<400> 131  
 aaggatcgat agcagttcat gatgtctaaa ctgggagatct tggaccat ctgtctgttt  
 60  
 ctgtttcccc ttactgctct tccgatggat ggagatcaat ctgcagaccg acttgcagag  
 120  
 cgtatgcagg acaacatttc atctgagcag catcccttct ttgaaaagag aggacgagac  
 180  
 tggatgcacac ctccgaggaa atgcagagac cgagcctgca aacctaacc ttgttgcgg  
 240  
 ggataagctg ttgatgacca actttgttat acggc 275

<210> 132  
 <211> 75  
 <212> PRT  
 <213> Conus lynceus

<400> 132  
 Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15

Pro Leu Thr Ala Leu Pro Met Asp Gly Asp Gln Ser Ala Asp Arg Leu

20

25

30

Ala Glu Arg Met Gln Asp Asn Ile Ser Ser Glu Gln His Pro Phe Phe  
 35 40 45

Glu Lys Arg Gly Arg Asp Cys Cys Thr Pro Pro Arg Lys Cys Arg Asp  
 50 55 60

Arg Ala Cys Lys Pro Gln Arg Cys Cys Gly Gly  
 65 70 75

&lt;210&gt; 133

&lt;211&gt; 23

&lt;212&gt; PRT

&lt;213&gt; Conus lynceus

&lt;220&gt;

&lt;221&gt; PEPTIDE

&lt;222&gt; (1)..(23)

&lt;223&gt; Xaa at residue 7, 8 and 18 is Pro or Hyp

&lt;400&gt; 133

Gly Arg Asp Cys Cys Thr Xaa Xaa Arg Lys Cys Arg Asp Arg Ala Cys  
 1 5 10 15

Lys Xaa Gln Arg Cys Cys Gly  
 20

&lt;210&gt; 134

&lt;211&gt; 803

&lt;212&gt; DNA

&lt;213&gt; Conus magus

&lt;400&gt; 134

caagagggtt cgatagcagt tcatgatgtc taaaactggga gtcttgttga ccatctgtct 60

gcttctgttt ccccttaactg ctcttccgat ggtggagat gaaacctgcaa accgacactgt 120

cgagcgtatg caggacaaca tttcatctga gcagttatccc ttgtttgaga agagacgaga 180

ttgttgcaact ccggcgaaga aatgcaaaga ccgacaatgc aaaccccaaga gatgttgcgc 240

tggacgataa cgttttgatg accaacttta tcacggctac gtcaagtgtt tagtgaataa 300

gtaaaatgtat tgcagtcttgc ttcagatttt cttttgtt ttggctaaa gatcaatgac 360

caaaccgttg ttttgcgtcg gattgtcata tatttctcgat ttccaatcca acactagatg 420

atttaaatcac gatagattaa ttttctatca atgccttgat ttttgcgtcg tcataatcagt 480

ttttgtttata tttttttttt cgtcaactgtc tacacaaacg catgcgtcg cgcgtgcacg 540

cacacacgca cgcacgctcg cacaaacatg cgcgcgcacg cacacacaca cacacacaca 600

caaacacaca cacgaagcaa tcacacaatt agttgacatt atttatattat tcattgtatgt 660

atttgttatt cgtttgcgtt tttttagaat agttgaggc cgtcttttg gatttatattt 720

aactgcattta ttgtatacga gtacttcgtt cggggaaaca ctgctgaaaa taaaacaaac 780

actgacgttag caaaaaaaaaaaa aaa 803

&lt;210&gt; 135

&lt;211&gt; 75

&lt;212&gt; PRT

&lt;213&gt; Conus magus

&lt;400&gt; 135

Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Phe  
1 5 10 15Pro Leu Thr Ala Leu Pro Met Asp Gly Asp Glu Pro Ala Asn Arg Pro  
20 25 30Val Glu Arg Met Gln Asp Asn Ile Ser Ser Glu Gln Tyr Pro Leu Phe  
35 40 45Glu Lys Arg Arg Asp Cys Cys Thr Pro Pro Lys Lys Cys Lys Asp Arg  
50 55 60Gln Cys Lys Pro Gln Arg Cys Cys Ala Gly Arg  
65 70 75

&lt;210&gt; 136

&lt;211&gt; 22

&lt;212&gt; PRT

&lt;213&gt; Conus magus

&lt;220&gt;

&lt;221&gt; PEPTIDE

&lt;222&gt; (1)..(22)

&lt;223&gt; Xaa at residue 6 and 7 is Pro or Hyp

&lt;400&gt; 136

Arg Asp Cys Cys Thr Xaa Xaa Lys Lys Cys Lys Asp Arg Gln Cys Lys  
1 5 10 15Xaa Gln Arg Cys Cys Ala  
20

&lt;210&gt; 137

&lt;211&gt; 656

&lt;212&gt; DNA

&lt;213&gt; Conus magus

&lt;400&gt; 137

caagagggtt cgatagcagt tcatgtatgtc taaaactggga gtcttgttga ccatctgtct 60

gcttctgttt ccccttaactg ctcttccaaat ggtggagat caacctgcag accaacacctgc 120

agatcgatcg caggacgaca tttcatctga gcagtatccc ttgtttgata tgagaaaaag 180

gtgttgcggc cccggcggtt catgccccgt atatttcaga gacaatttttta ttgttgggtt 240

ttgtttaatg acaacgtgtc gatgaccaac ttcattatca cgactacgcc aagtgtctaa 300

tgaataaaata aaatgtttgc agtctcgctc agatttgtt ttgttatttttgc tctaaagat 360

caatgaccaa accgttggttt tgggtgtggat tttcatatat ttctcgagtc ctatccaaca 420

ctagatgatt taatcacgat agatctgatt tttttatcaa aggcttggttt ttctgtctgt 480

cacatcagtt ttgttataat ttaatttttc gtcactgatt acacacacgc atgaacgcac 540

agagtactaa cacatacaca cacacacaca cacacacaca cacacacaca cacacacaca 600

cacacacaca cacgcgcgcg cgccgcgcgc tctagtagcg ccgcgcac acacac 656

&lt;210&gt; 138

&lt;211&gt; 74

&lt;212&gt; PRT

&lt;213&gt; Conus magus

&lt;400&gt; 138

Met	Met	Ser	Lys	Leu	Gly	Val	Leu	Leu	Thr	Ile	Cys	Leu	Leu	Leu	Phe
1				5				10					15		

Pro	Leu	Thr	Ala	Leu	Pro	Met	Asp	Gly	Asp	Gln	Pro	Ala	Asp	Gln	Pro
	20				25						30				

Ala	Asp	Arg	Met	Gln	Asp	Asp	Ile	Ser	Ser	Glu	Gln	Tyr	Pro	Leu	Phe
	35				40					45					

Asp	Met	Arg	Lys	Arg	Cys	Cys	Gly	Pro	Gly	Gly	Ser	Cys	Pro	Val	Tyr
	50				55				60						

Phe	Arg	Asp	Asn	Phe	Ile	Cys	Gly	Cys	Cys						
	65				70										

&lt;210&gt; 139

&lt;211&gt; 21

&lt;212&gt; PRT

&lt;213&gt; Conus magus

&lt;220&gt;

&lt;221&gt; PEPTIDE

&lt;222&gt; (1)..(21)

<223> Xaa at residue 4 and 9 is Pro or Hyp; Xaa at residue is 11 Tyr, 1 25I-Tyr, mono-iodo-Tyr, di-iodo-Tyr, O-sulpho-Tyr or O-phospho-Ty

&lt;400&gt; 139

Cys	Cys	Gly	Xaa	Gly	Gly	Ser	Cys	Xaa	Val	Xaa	Phe	Arg	Asp	Asn	Phe
1				5					10				15		

Ile	Cys	Gly	Cys	Cys											
	20														

&lt;210&gt; 140

&lt;211&gt; 594

&lt;212&gt; DNA

&lt;213&gt; Conus magus

&lt;400&gt; 140

caagaggat cgatacgagt tcatgatgtc taaaactggga gtcttgtga ccatctgttt 60

gcttctgttt ccccttactg ctcttccgag ggatggagat caatctgttag accgacactgc 120

agagcgtatg caggacgaca tttcatctga gctgcattcc ttgtcaatca gaaaaaagaat 180

gtgttgcggc gagagtgcgc catgccccca gatattcaga aacagtcaga ttgtcattg 240

ttgttaatg acaacgtgtc gatgaccacc ttcgttatca cgactaatga taagtaaaat 300

gattgcagtc tcgctcagat ttgctttgt atttggct aagatcaat gaccaaaccg 360

ttgtttgtat gtggattttc atatatttct cgagtcctat ccaacactag atgatttaat 420

cacgatagat ctgattttt tatcaaagcc ttggttttc gtctgtcaca tcagtttgc 480

ttatatttaa ttttcgtca ctgattacac acacgcgtca acgcacagac gtactaacac 540

atacacacac acacacacac acacacacac acacacacac acac 594

&lt;210&gt; 141

&lt;211&gt; 74

<212> PRT  
 <213> Conus magus

<400> 141  
 Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15

Pro Leu Thr Ala Leu Pro Arg Asp Gly Asp Gln Ser Val Asp Arg Pro  
 20 25 30

Ala Glu Arg Met Gln Asp Asp Ile Ser Ser Glu Leu His Pro Leu Ser  
 35 40 45

Ile Arg Lys Arg Met Cys Cys Gly Glu Ser Ala Pro Cys Pro Ser Tyr  
 50 55 60

Phe Arg Asn Ser Gln Ile Cys His Cys Cys  
 65 70

<210> 142  
 <211> 22  
 <212> PRT  
 <213> Conus magus

<220>  
 <221> PEPTIDE  
 <222> (1)..(22)  
 <223> Xaa at residue 5 is Glu or gamma-carboxy Glu; Xaa at residue 8 and 10 is Pro or Hyp; Xaa at residue 12 is Tyr, 125I-Tyr, mono-iodo-Tyr, di-iodo-Tyr, O-sulpho-Tyr or O-phospho-Ty

<400> 142  
 Met Cys Cys Gly Xaa Ser Ala Xaa Cys Xaa Ser Xaa Phe Arg Asn Ser  
 1 5 10 15

Gln Ile Cys His Cys Cys  
 20

<210> 143  
 <211> 501  
 <212> DNA  
 <213> Conus magus

<400> 143  
 caagagggtt cgatagcagt tcatgtatgtc taaaactggga gtcttggat ccatctgtct 60  
 gcttctgttt ccccttactg ctcttccaaat ggatggagat caacctgcag accaacctgc 120  
 agatcgatgtc caggacgaca tttcatctgtc gcagttatccc ttgtttgata agagacaaaa 180  
 gtgttgcggc cccggcggtt catggccgtt atatccaca gacaatttta ttgttgggtt 240  
 ttgtttaatgtt acaacgtgtc gatgaccaac ttcattatca cgactacgcc aagtgtctaa 300  
 tgaataaataa aaatgtttttt agtctcgctc agatggctt ttgttattttgg tctaaagatc 360  
 aatgaccaaa ccgttgggtt ggtgctggat tttcatatat ttctcgattc ctatccaaca 420  
 ctagatgatt taatcacat agatctgatt ttttatcaa tgccttaatt ttttgctctg 480  
 tcataatcagt tttgtttata t 501

<210> 144  
 <211> 74  
 <212> PRT

&lt;213&gt; Conus magus

<400> 144  
 Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Phe  
 1 5 10 15  
 Pro Leu Thr Ala Leu Pro Met Asp Gly Asp Gln Pro Ala Asp Gln Pro  
 20 25 30  
 Ala Asp Arg Met Gln Asp Asp Ile Ser Ser Glu Gln Tyr Pro Leu Phe  
 35 40 45  
 Asp Lys Arg Gln Lys Cys Cys Gly Pro Gly Gly Ser Cys Pro Val Tyr  
 50 55 60  
 Phe Thr Asp Asn Phe Ile Cys Gly Cys Cys  
 65 70

<210> 145  
 <211> 23  
 <212> PRT  
 <213> Conus magus

<220>  
 <221> PEPTIDE  
 <222> (1)..(23)  
 <223> Xaa at residue 1 is Gln or pyro-Glu; Xaa at residue 6 and 11 is P  
 ro or Hyp; Xaa at residue 13 is Tyr, 125I-Tyr, mono-iodo-Tyr, di-  
 iodo-Tyr, O-sulpho-Tyr or O-phospho-Ty

<400> 145  
 Xaa Lys Cys Cys Gly Xaa Gly Gly Ser Cys Xaa Val Xaa Phe Thr Asp  
 1 5 10 15  
 Asn Phe Ile Cys Gly Cys Cys  
 20

<210> 146  
 <211> 454  
 <212> DNA  
 <213> Conus magus

<400> 146  
 caagaggat cgatagcagt tcatgatgtc taaaactggga gtcttggta ccatctgtct 60  
 gcttctgttt ccccttactg ctcttccaaat ggatggagat caacctgcag accaacctgc 120  
 agatcgatg caggacgaca tttcatctga gcagtatccc ttgtttgata agagacaaaa 180  
 gtgttgcggc cccggcgggtt catgccccgt atatttcaga gacaatttta ttgttggttg 240  
 ttgttaaatg acaacgtgtc gatgaccatc ttcattatca cgactacgcc aagtgtctaa 300  
 tgaataaataa aatgattgc agtctcgctc agatttgttt ttgttatttg gtctaaagat 360  
 caatgaccaa accgttgggtt tgggtggat tttcatatat ttctcgattc ctatccaaca 420  
 ctagatgatt taatcacgat agatctgatt tttt 454

<210> 147  
 <211> 74  
 <212> PRT  
 <213> Conus magus

&lt;400&gt; 147

Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15

Pro Leu Thr Ala Leu Pro Met Asp Gly Asp Gln Pro Ala Asp Gln Pro  
 20 25 30

Ala Asp Arg Met Gln Asp Asp Ile Ser Ser Glu Gln Tyr Pro Leu Phe  
 35 40 45

Asp Lys Arg Gln Lys Cys Cys Gly Pro Gly Gly Ser Cys Pro Val Tyr  
 50 55 60

Phe Arg Asp Asn Phe Ile Cys Gly Cys Cys  
 65 70

<210> 148

<211> 23

<212> PRT

<213> Conus magus

<220>

<221> PEPTIDE

<222> (1)..(23)

<223> Xaa at residue 1 is Gln or pyro-Glu; Xaa at residue 6 and 11 is P  
 ro or Hyp; Xaa at residue 13 is Tyr, 125I-Tyr, mono-iodo-Tyr, di-  
 iodo-Tyr, O-sulpho-Tyr or O-phospho-Ty

<400> 148

Xaa Lys Cys Cys Gly Xaa Gly Gly Ser Cys Xaa Val Xaa Phe Arg Asp  
 1 5 10 15

Asn Phe Ile Cys Gly Cys Cys  
 20

<210> 149

<211> 22

<212> PRT

<213> Conus magus

<220>

<221> PEPTIDE

<222> (1)..(22)

<223> Xaa at residue 1 is Gln or pyro-Glu; Xaa at residue 10 and 20 is  
 Pro or Hyp; Xaa at residue 12 is Tyr, 125I-Tyr, mono-iodo-Tyr, di-  
 iodo-Tyr, O-sulpho-Tyr or O-phospho-Ty

<400> 149

Xaa Lys Cys Cys Ser Gly Gly Ser Cys Xaa Leu Xaa Phe Arg Asp Arg  
 1 5 10 15

Leu Ile Cys Xaa Cys Cys  
 20

<210> 150

<211> 19

<212> PRT

<213> Conus marmoreus

<220>

<221> PEPTIDE

<222> (1)..(19)

<223> Xaa at residue 16 is Pro or Hyp

<400> 150

Ser Lys Gln Cys Cys His Leu Ala Ala Cys Arg. Phe Gly Cys Thr Xaa  
 1 5 10 15

Cys Cys Asn

<210> 151  
 <211> 321  
 <212> DNA  
 <213> Conus marmoreus  
 <400> 151  
 caagaaggat cgatagcagt tcatgatgtc taaaactggga gtcttgttga ccatctgtct 60  
 gcttctgttt cccgttactg ctcttccgat ggatggtgat caacctgcag accgacttgt 120  
 agagcgtatg caggacaaca tttcatctga gcagcatccc ttctttgaaa agagaagagg 180  
 aggctgttgc acacccctccga ggaaatgcaa agaccgagcc tgcaaacctg cacgttgctg 240  
 cggcccagga taacgtgttgc atgaccaact ttgttatcac ggctacgtca agtgtctagt 300  
 gaataagtaa aacgattgca g 321  
  
 <210> 152  
 <211> 76  
 <212> PRT  
 <213> Conus marmoreus  
  
 <400> 152  
 Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe 1 5 10 15  
 Pro Val Thr Ala Leu Pro Met Asp Gly Asp Gln Pro Ala Asp Arg Leu 20 25 30  
 Val Glu Arg Met Gln Asp Asn Ile Ser Ser Glu Gln His Pro Phe Phe 35 40 45  
 Glu Lys Arg Arg Gly Gly Cys Cys Thr Pro Pro Arg Lys Cys Lys Asp 50 55 60  
 Arg Ala Cys Lys Pro Ala Arg Cys Cys Gly Pro Gly 65 70 75  
  
 <210> 153  
 <211> 24  
 <212> PRT  
 <213> Conus marmoreus  
  
 <220>  
 <221> PEPTIDE  
 <222> (1)...(24)  
 <223> Xaa at residue 3, 8, 18 and 24 is Pro or Hyp  
  
 <400> 153  
 Arg Gly Gly Cys Cys Thr Xaa Xaa Arg Lys Cys Lys Asp Arg Ala Cys 1 5 10 15  
 Lys Xaa Ala Arg Cys Cys Gly Xaa 20  
  
 <210> 154  
 <211> 296  
 <212> DNA  
 <213> Conus marmoreus  
  
 <400> 154  
 gagctcggtt ccccgacctc aagagggatc gatagcagt catgtatgtct aaactggaa 60

tctttgttgcacatctgtctatcttcccttactgc tgttccgctg gatggagatc 120  
aacctgcaga ccgacccgtcagagcgtatgc aggacgacat ttcatctgaa catcatccct 180  
tttttgcattcgtaaaacgg tggttcaggt tatcatgcgg cctggatgc cacccttgc 240  
gtggatgacc agctttgtta tcgcggccctc atcaagtgtc taatgaataa gtaaaa 296  
<210> 155  
<211> 68  
<212> PRT  
<213> Conus marmoreus  
<400> 155  
Met Met Ser Lys Leu Gly Ile Leu Leu Thr Ile Cys Leu Leu Leu Phe 1 15  
1 5 10 15  
Pro Leu Thr Ala Val Pro Leu Asp Gly Asp Gln Pro Ala Asp Arg Pro 20 25 30  
Ala Glu Arg Met Gln Asp Asp Ile Ser Ser Glu His His Pro Phe Phe 35 40 45  
Asp Pro Val Lys Arg Cys Cys Arg Leu Ser Cys Gly Leu Gly Cys His 50 55 60  
Pro Cys Cys Gly  
65  
<210> 156  
<211> 14  
<212> PRT  
<213> Conus marmoreus  
<220>  
<221> PEPTIDE  
<222> (1)..(14)  
<223> Xaa at residue 12 is Pro or Hyp  
<400> 156  
Cys Cys Arg Leu Ser Cys Gly Leu Gly Cys His Xaa Cys Cys 1 5 10  
<210> 157  
<211> 355  
<212> DNA  
<213> Conus marmoreus  
<400> 157  
ggcctacacc aagcttgcattgcctgcaggt cgactctaga ggatccccga tcgatagcag 60  
ttcatgtatgttctagactggg agtcttgttg accatctgtc tacttctgtt tcccccttact 120  
gctgttccgc tggatggaga tcaacctgcg gaccgacctg cagagcgcct gcaggagc 180  
atttcatctg aacatcatcc ccattttgtat tccggcagag agtgttgcgg ttcttcgc 240  
tgccgccttg gatgcgtgcc ttgttgcgttgat tgaccagtt tgttatcacg gcctcatcga 300  
gtgtctaatg aataagtaaa acgattgcag taggcggta ccgagctcga attcc 355  
<210> 158  
<211> 69  
<212> PRT

&lt;213&gt; Conus marmoreus

&lt;400&gt; 158

Met Met Ser Arg Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
1 5 10 15Pro Leu Thr Ala Val Pro Leu Asp Gly Asp Gln Pro Ala Asp Arg Pro  
20 25 30Ala Glu Arg Leu Gln Asp Asp Ile Ser Ser Glu His His Pro His Phe  
35 40 45Asp Ser Gly Arg Glu Cys Cys Gly Ser Phe Ala Cys Arg Phe Gly Cys  
50 55 60Val Pro Cys Cys Val  
65

&lt;210&gt; 159

&lt;211&gt; 17

&lt;212&gt; PRT

&lt;213&gt; Conus marmoreus

&lt;220&gt;

&lt;221&gt; PEPTIDE

&lt;222&gt; (1)..(17)

&lt;223&gt; Xaa at residue 1 is Glu or gamma-carboxy Glu; Xaa at residue 14 is Pro or Hy

&lt;400&gt; 159

Xaa Cys Cys Gly Ser Phe Ala Cys Arg Phe Gly Cys Val Xaa Cys Cys  
1 5 10 15

Val

&lt;210&gt; 160

&lt;211&gt; 295

&lt;212&gt; DNA

&lt;213&gt; Conus marmoreus

&lt;400&gt; 160

cgacacctaa agggatcgat agcagttcat gatgtctaaa ctgggagtct tggaccat 60

ctgtctactt ctatcccc ttactgctgt tccgctggat ggagaccaac ctgcagaccg 120

acctgcagag cgtatgcagg acgacatttc atctgaacgt catcctttt ttgatcgac 180

caaacagtgt tgccatctgc cggatgccg cttcgatgt acgcctgtt gttggatgc 240

agctttgtta tcgcgtcctc atcaagtgtc taatgaataa gtaaaatgtat tgca 295

&lt;210&gt; 161

&lt;211&gt; 67

&lt;212&gt; PRT

&lt;213&gt; Conus marmoreus

&lt;400&gt; 161

Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
1 5 10 15Pro Leu Thr Ala Val Pro Leu Asp Gly Asp Gln Pro Ala Asp Arg Pro  
20 25 30Ala Glu Arg Met Gln Asp Asp Ile Ser Ser His Pro Phe Phe Asp Arg  
35 40 45

Ser Lys Gln Cys Cys His Leu Pro Ala Cys Arg Phe Gly Cys Thr Pro  
 50 55 60

Cys Cys Trp  
 65

<210> 162

<211> 19

<212> PRT

<213> Conus marmoreus

<220>

<221> PEPTIDE

<222> (1)..(19)

<223> Xaa at residue 8 and 16 is Pro or Hyp; Xaa at residue 19 is Trp or bromo-Tr

<400> 162

Ser Lys Gln Cys Cys His Leu Xaa Ala Cys Arg Phe Gly Cys Thr Xaa  
 1 5 10 15

Cys Cys Xaa

<210> 163

<211> 235

<212> DNA

<213> Conus marmoreus

<400> 163

ggatccatga tgtctaaact gggagtcttg ttgaccatct gtctgcttct gtttcccctt 60

actgctcttc cgctggatgg agatcaacct gcagaccaac gtgcagagcg tacgcaggcc 120

gagaagcatt cttgcctga tccgagaatg ggctgttgcc cgttccatg caaaaccagt 180

tgcactactt tgtgttgccg gtgtatgataa cgtgttgatg accaactttc tcgag 235

<210> 164

<211> 67

<212> PRT

<213> Conus marmoreus

<400> 164

Gly Ser Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu  
 1 5 10 15

Leu Phe Pro Leu Thr Ala Leu Pro Leu Asp Gly Asp Gln Pro Ala Asp  
 20 25 30

Gln Arg Ala Glu Arg Thr Gln Ala Glu Lys His Ser Leu Pro Asp Pro  
 35 40 45

Arg Met Gly Cys Cys Pro Phe Pro Cys Lys Thr Ser Cys Thr Thr Leu  
 50 55 60

Cys Cys Gly  
 65

<210> 165

<211> 17

<212> PRT

<213> Conus marmoreus

<220>

<221> PEPTIDE  
 <222> (1)..(17)  
 <223> Xaa at residue 5 and 7 is Pro or Hyp

<400> 165  
 Met Gly Cys Cys Xaa Phe Xaa Cys Lys Thr Ser Cys Thr Thr Leu Cys  
 1 5 10 15

Cys

<210> 166  
 <211> 16  
 <212> PRT  
 <213> Conus marmoreus

<220>  
 <221> PEPTIDE  
 <222> (1)..(16)  
 <223> Xaa at residue 4 and 6 is Trp or bromo-Trp

<400> 166  
 Cys Cys His Xaa Asn Xaa Cys Asp His Leu Cys Ser Cys Cys Gly Ser  
 1 5 10 15

<210> 167  
 <211> 357  
 <212> DNA  
 <213> Conus marmoreus

<400> 167  
 gccaagttg catgcctgca ggatgactct agaggatccc cacctcaaga gggatcgata 60  
 gcagttcatg atgtctaaac tgggagtc ttgtgaccatc tgtctacttc tgtttgcct 120  
 tactgctgtt ccgctggatg gagatcaacc tgcagaccga cctgcagaac gtatgcagga 180  
 cgacatttca tctgaacgatc atcccatgtt tgatgccgtc agagattgtt gcccgttgc 240  
 ggcatgcccc ttggatgca acccttgttg tggatgacca gctttgttat cgggacctca 300  
 tcaagtgtct aatgaataag taaaaaacgat ttcgagtttgg taccgagctc gaattcc 357

<210> 168  
 <211> 67  
 <212> PRT  
 <213> Conus marmoreus

<400> 168  
 Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15

Ala Leu Thr Ala Val Pro Leu Asp Gly Asp Gln Pro Ala Asp Arg Pro  
 20 25 30

Ala Glu Arg Met Gln Asp Asp Ile Ser Ser His Pro Met Phe Asp Ala  
 35 40 45

Val Arg Asp Cys Cys Pro Leu Pro Ala Cys Pro Phe Gly Cys Asn Pro  
 50 55 60

Cys Cys Gly  
 65

<210> 169  
 <211> 16

<212> PRT  
<213> Conus marmoreus

<220>  
<221> PEPTIDE  
<222> (1)..(16)  
<223> Xaa at residue 4, 6, 9 and 14 is Pro or Hyp

<400> 169  
Asp Cys Cys Xaa Leu Xaa Ala Cys Xaa Phe Gly Cys Asn Xaa Cys Cys  
1 5 10 15

<210> 170  
<211> 16  
<212> PRT  
<213> Conus marmoreus

<220>  
<221> PEPTIDE  
<222> (1)..(16)  
<223> Xaa at residue 4 and 13 is Pro or Hyp

<400> 170  
Cys Cys Ala Xaa Ser Ala Cys Arg Leu Gly Cys Arg Xaa Cys Cys Arg  
1 5 10 15

<210> 171  
<211> 16  
<212> PRT  
<213> Conus marmoreus

<220>  
<221> PEPTIDE  
<222> (1)..(16)  
<223> Xaa at residue 4 and 13 is Pro or Hyp

<400> 171  
Cys Cys Ala Xaa Ser Ala Cys Arg Leu Gly Cys Arg Xaa Cys Cys Arg  
1 5 10 15

<210> 172  
<211> 16  
<212> PRT  
<213> Conus marmoreus

<220>  
<221> PEPTIDE  
<222> (1)..(16)  
<223> Xaa at residue 4 and 13 is Pro or Hyp

<400> 172  
Cys Cys Ala Xaa Ser Ala Cys Arg Leu Gly Cys Arg Xaa Cys Cys Arg  
1 5 10 15

<210> 173  
<211> 17  
<212> PRT  
<213> Conus marmoreus

<220>  
<221> PEPTIDE  
<222> (1)..(17)  
<223> Xaa at residue 14 is Pro or Hyp

<400> 173

51

Gly	Cys	Cys	Gly	Ser	Phe	Ala	Cys	Arg	Phe	Gly	Cys	Val	Xaa	Cys	Cys
1				5				10					15		

Val

<210> 174  
<211> 244  
<212> DNA  
<213> *Conus nobilis*

<400> 174  
ggatccatga tgtctaaact gggagtcttg ttgaccatct gtctacttct gtttccccctt 60  
actgctcttc cgctggatga agatcaacccg gtacaccgac ctgcagagcg tatgcaggac 120  
atttcatctg atcaacatct cttctttgat ctcataaac ggtgctgcga gttgccatgc 180  
gggccagggct tttgcgtccc ttgttgctga catcaataac gtgttgatga ccaactttct 240  
cgag 244

<210> 175  
<211> 69  
<212> PRT  
<213> *Conus nobilis*

<400> 175  
Gly Ser Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu  
1 5 10 15

Leu Phe Pro Leu Thr Ala Leu Pro Leu Asp Glu Asp Gln Pro Val His  
20 25 30

Arg Pro Ala Glu Arg Met Gln Asp Ile Ser Ser Asp Gln His Leu Phe  
35 40 45

Phe Asp Leu Ile Lys Arg Cys Cys Glu Leu Pro Cys Gly Pro Gly Phe  
50 55 60

Cys Val Pro Cys Cys  
65

<210> 176  
<211> 15  
<212> PRT  
<213> *Conus nobilis*

<220>  
<221> PEPTIDE  
<222> (1)..(15)  
<223> Xaa at residue 3 is Glu or gamma-carboxy Glu; Xaa at residue 5, 8  
adn 13 is Pro or Hy

<400> 176  
Cys Cys Xaa Leu Xaa Cys Gly Xaa Gly Phe Cys Val Xaa Cys Cys  
1 5 10 15

<210> 177  
<211> 262  
<212> DNA  
<213> *Conus nobilis*

<400> 177  
ggatccatga tgtctaaact gggagtcttg ttgaccatct gtctacttct gtttccccctt 60

actgcttttc cgatggatgg agatcaacct gcagaccaac ctgcagatcg tatgcaggac 120  
 gacatttcat ctgagcagta tcccttgttt gataagagac aaaagtgtt cactggaaag 180  
 aaggggatcat gctccggcaa agcatgcaaa aatctcaaat gttgctctgg acgataacgt 240  
 gttgatgacc aactttctcg ag 262  
  
 <210> 178  
 <211> 78  
 <212> PRT  
 <213> Conus nobilis  
  
 <400> 178  
 Gly Ser Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu  
 1 5 10 15  
  
 Leu Phe Pro Leu Thr Ala Phe Pro Met Asp Gly Asp Gln Pro Ala Asp  
 20 25 30  
  
 Gln Pro Ala Asp Arg Met Gln Asp Asp Ile Ser Ser Glu Gln Tyr Pro  
 35 40 45  
  
 Leu Phe Asp Lys Arg Gln Lys Cys Cys Thr Gly Lys Lys Gly Ser Cys  
 50 55 60  
  
 Ser Gly Lys Ala Cys Lys Asn Leu Lys Cys Cys Ser Gly Arg  
 65 70 75  
  
 <210> 179  
 <211> 23  
 <212> PRT  
 <213> Conus nobilis  
  
 <220>  
 <221> PEPTIDE  
 <222> (1)...(23)  
 <223> Xaa at residue 1 is Gln or pyro-Glu  
  
 <400> 179  
 Xaa Lys Cys Cys Thr Gly Lys Lys Gly Ser Cys Ser Gly Lys Ala Cys  
 1 5 10 15  
  
 Lys Asn Leu Lys Cys Cys Ser  
 20  
  
 <210> 180  
 <211> 238  
 <212> DNA  
 <213> Conus pulicarius  
  
 <400> 180  
 ggatccatga tgtctaaact gggagttttt ttgaccatct gtctgcttct gtttccccctt 60  
 actgctgttc cgctggatgg agatcaacct gcagacccgac ctgcagagcg tatgcaggac 120  
 attgcaactg aacagcatcc cttctttgat cccgtcaaac ggtgttgcaa cagctgttac 180  
 atgggatgca tcccttgtt cttcttagtaa taacgtgtt atgaccaact ttctcgag 238  
  
 <210> 181  
 <211> 68  
 <212> PRT  
 <213> Conus pulicarius

<400> 181  
 Gly Ser Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu  
 1 5 10 15

Leu Phe Pro Leu Thr Ala Val Pro Leu Asp Gly Asp Gln Pro Ala Asp  
 20 25 30

Arg Pro Ala Glu Arg Met Gln Asp Ile Ala Thr Glu Gln His Pro Phe  
 35 40 45

Phe Asp Pro Val Lys Arg Cys Cys Asn Ser Cys Tyr Met Gly Cys Ile  
 50 55 60

Pro Cys Cys Phe  
 65

<210> 182

<211> 14

<212> PRT

<213> Conus pulicarius

<220>

<221> PEPTIDE

<222> (1)...(14)

<223> Xaa at residue 11 is Pro or Hyp; Xaa at residue 5 is Tyr, 125I-Ty  
 r, mono-iodo-Tyr, di-iodo-Tyr, O-sulpho-Tyr or O-phospho-Ty

<400> 182

Cys Cys Asn Ser Cys Xaa Met Gly Cys Ile Xaa Cys Cys Phe  
 1 5 10

<210> 183

<211> 238

<212> DNA

<213> Conus quercinus

<400> 183

ggatccatga tgtctaaact gggagtcttg ttgaccatct gtctgcttct gtttccctt 60

acagctcttc agctggatgg agatcaacct gcagaccgac ctgcagagcg tacgcaggac 120

attgcatctg aacagtatcg aaagtttgat cagagacaga ggtgttgcca gtggccatgc 180

cccggtagtt gcagatgctg ccgtactggt taacgtttg atgaccaact ttctcgag 238

<210> 184

<211> 70

<212> PRT

<213> Conus quercinus

<400> 184

Gly Ser Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu  
 1 5 10 15

Leu Phe Pro Leu Thr Ala Leu Gln Leu Asp Gly Asp Gln Pro Ala Asp  
 20 25 30

Arg Pro Ala Glu Arg Thr Gln Asp Ile Ala Ser Glu Gln Tyr Arg Lys  
 35 40 45

Phe Asp Gln Arg Gln Arg Cys Cys Gln Trp Pro Cys Pro Gly Ser Cys  
 50 55 60

Arg Cys Cys Arg Thr Gly  
 65 70

<210> 185  
 <211> 17  
 <212> PRT  
 <213> Conus quercinus

<220>  
 <221> PEPTIDE  
 <222> (1)..(17)  
 <223> Xaa at residue 1 is Gln or pyro-Glu; Xaa at residue 7 and 9 is Pro or Hyp; Xaa at residue 6 is Trp or bromo-Tr

<400> 185  
 Xaa Arg Cys Cys Gln Xaa Xaa Cys Xaa Gly Ser Cys Arg Cys Cys Arg  
 1 5 10 15

Thr

<210> 186  
 <211> 15  
 <212> PRT  
 <213> Conus quercinus

<220>  
 <221> PEPTIDE  
 <222> (1)..(15)  
 <223> Xaa at residue 11 and 14 is Pro or Hyp

<400> 186  
 Cys Cys Ser Gln Asp Cys Leu Val Cys Ile Xaa Cys Cys Xaa Asn  
 1 5 10 15

<210> 187  
 <211> 15  
 <212> PRT  
 <213> Conus quercinus

<220>  
 <221> PEPTIDE  
 <222> (1)..(15)  
 <223> Xaa at residue 11 14 is Pro or Hyp; Xaa at residue 7 is Trp or bromo-Tr

<400> 187  
 Cys Cys Ser Arg His Cys Xaa Val Cys Ile Xaa Cys Cys Xaa Asn  
 1 5 10 15

<210> 188  
 <211> 323  
 <212> DNA  
 <213> Conus radiatus

<400> 188  
 tcaagaagga tcgatagcag ttcatgatgt ctaaactggg agtcttggc accatctgtc 60  
 tgcttcgtt tcccttact gctttccga tggatggaga tcaacctgtt gaccgacttg 120  
 cagagcgtat gcaggacaac atttcatctg agcagcatac cttcttgaa aagagactac 180  
 catcgtgtt ctccttaac ttgcggctt gcccagtacc agcatgaaa cgtaaccctt 240  
 gttgcacagg ataacgtgtt gatgaccaac tttgttatca cggctacgtc aagtgtctag 300  
 tgaataagta aaacgattgc agt 323

<210> 189  
 <211> 76  
 <212> PRT  
 <213> Conus radiatus

<400> 189  
 Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Phe  
 1 5 10 15

Pro Leu Thr Ala Leu Pro Met Asp Gly Asp Gln Pro Val Asp Arg Leu  
 20 25 30

Ala Glu Arg Met Gln Asp Asn Ile Ser Ser Glu Gln His Thr Phe Phe  
 35 40 45

Glu Lys Arg Leu Pro Ser Cys Cys Ser Leu Asn Leu Arg Leu Cys Pro  
 50 55 60

Val Pro Ala Cys Lys Arg Asn Pro Cys Cys Thr Gly  
 65 70 75

<210> 190  
 <211> 24  
 <212> PRT  
 <213> Conus radiatus

<220>  
 <221> PEPTIDE  
 <222> (1)..(24)  
 <223> Xaa at residue 2, 13, 15 and 21 is Pro or Hyp

<400> 190  
 Leu Xaa Ser Cys Cys Ser Leu Asn Leu Arg Leu Cys Xaa Val Xaa Ala  
 1 5 10 15

Cys Lys Arg Asn Xaa Cys Cys Thr  
 20

<210> 191  
 <211> 336  
 <212> DNA  
 <213> Conus radiatus

<400> 191  
 aggtcgactc tagaggatcc ccaaggatcg atagcagttc atgatgtcta aactgggagt 60  
 cttgttgacc atctgtctgc ttctgtttcc ctttactgtc cttccgatgg atggagatca 120  
 acctgcagac cgacttgcag agcgtatgca ggacgacatt tcatctgagc agcatccctt 180  
 cttaaaaaag agacaacaaa gatgttgac cgttaagagg atttgtccag taccagcatg 240  
 cagaagtaaa ctttggca aatcataacg tattgtgac caactttgtt atcacggcta 300  
 cgtcaagtgt ctatgtata agtaaaatga ttgcag 336

<210> 192  
 <211> 75  
 <212> PRT  
 <213> Conus radiatus

<400> 192  
 Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Phe  
 1 5 10 15

56

Pro Leu Thr Ala Leu Pro Met Asp Gly Asp Gln Pro Ala Asp Arg Leu  
 20 25 30

Ala Glu Arg Met Gln Asp Asp Ile Ser Ser Glu Gln His Pro Phe Phe  
 35 40 45

Lys Lys Arg Gln Gln Arg Cys Cys Thr Val Lys Arg Ile Cys Pro Val  
 50 55 60

Pro Ala Cys Arg Ser Lys Pro Cys Cys Lys Ser  
 65 70 75

<210> 193

<211> 24

<212> PRT

<213> Conus radiatus

<220>

<221> PEPTIDE

<222> (1)...(24)

<223> Xaa at residue 1 is Gln or pyro-Glu; Xaa at residue 12, 14 and 20  
 is Pro or Hy

<400> 193

Xaa Gln Arg Cys Cys Thr Val Lys Arg Ile Cys Xaa Val Xaa Ala Cys  
 1 5 10 15

Arg Ser Lys Xaa Cys Cys Lys Ser  
 20

<210> 194

<211> 326

<212> DNA

<213> Conus radiatus

<400> 194

acctcaagaa ggatcgatag cagttcatga tgtctaaact gggagtcttg ttgaccatct 60

gtctgcttct gtttcccggtt actgctttc cgatggatgg tcatcaacct gcagaccgac 120

ttgttagagcg tatgcaggac aacatttcat ctgagcagca tcccttcttt gaaaagagaa 180

gaggaggctg ttgcacaccc ccgaggaaat gcaaagaccg agcctgcaaa cctgcacgtt 240

gctgcggccc aggataacgt gttgatgacc aactttgtta tcacggctac gtcaagtgtc 300

tagtgaataa gtaaaacgt tgca 326

<210> 195

<211> 76

<212> PRT

<213> Conus radiatus

<400> 195

Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15

Pro Val Thr Ala Leu Pro Met Asp Gly Asp Gln Pro Ala Asp Arg Leu  
 20 25 30

Val Glu Arg Met Gln Asp Asn Ile Ser Ser Glu Gln His Pro Phe Phe  
 35 40 45

Glu Lys Arg Arg Gly Gly Cys Cys Thr Pro Pro Arg Lys Cys Lys Asp  
 50 55 60

Arg Ala Cys Lys Pro Ala Arg Cys Cys Gly Pro Gly  
65 70 75

<210> 196  
<211> 24  
<212> PRT  
<213> *Conus radiatus*

<220>  
<221> PEPTIDE  
<222> (1)..(24)  
<223> Xaa at residue 7, 8, 18 and 24 is Pro or Hyp

<400> 196  
Arg Gly Gly Cys Cys Thr Xaa Xaa Arg Lys Cys Lys Asp Arg Ala Cys  
1 5 10 15

Lys Xaa Ala Arg Cys Cys Gly Xaa  
20

<210> 197  
<211> 238  
<212> DNA  
<213> *Conus rattus*

<400> 197  
ggatccatga tgtctaaact gggagtcgg gtgaccatct gcctgcttct gttccctctt 60  
gctgctttc cactggatgg agatcaacct gcagaccacc ctgcaaagcg tacgcaagat 120  
gacagttcag ctgcctgtat caatgcctgg cttgatgaat cccagacttg ctgcagtaac 180  
tgcggtaag attgtatgg ttgttgcag taacgtttg atgaccaact ttctcgag 238

<210> 198  
<211> 70  
<212> PRT  
<213> *Conus rattus*

<400> 198  
Gly Ser Met Met Ser Lys Leu Gly Val Leu Val Thr Ile Cys Leu Leu  
1 5 10 15

Leu Phe Pro Leu Ala Ala Phe Pro Leu Asp Gly Asp Gln Pro Ala Asp  
20 25 30

His Pro Ala Lys Arg Thr Gln Asp Asp Ser Ser Ala Ala Leu Ile Asn  
35 40 45

Ala Trp Leu Asp Glu Ser Gln Thr Cys, Cys Ser Asn Cys Gly Glu Asp  
50 55 60

Cys Asp Gly Cys Cys Gln  
65 70

<210> 199  
<211> 16  
<212> PRT  
<213> *Conus rattus*

<220>  
<221> PEPTIDE  
<222> (1)..(16)  
<223> Xaa at residue 1 is Gln or pyro-Glu; Xaa at residue 9 is Glu or gamma-carboxy Gl

<400> 199  
 Xaa Thr Cys Cys Ser Asn Cys Gly Xaa Asp Cys Asp Gly Cys Cys Gln  
 1 5 10 15

<210> 200

<211> 327

<212> DNA

<213> Conus stercusmuscarum

<400> 200

gacctcaaga gggatcgata gcagttcgtg atgtataaac tgggagtctt gttgaccatc 60  
 tgtctgttc tgttccctct tactgtcttt ccgatggatg gagatcaacc tgcagaccaa 120  
 cctgcagatc gtatgcagga cgacatttca tctgagcagt atcccttggt tgataagaga 180  
 caaaaagtgtt gcactggaa gaaggggtca tgctccggca aagcatgcaa aaatctcaaa 240  
 tgttgctctg gacgataacg tgttgatgac caactttgtt atcacggcta cgtcaagtgt 300  
 ctaatgaata agtaaaaacgta ttgcagt 327

<210> 201

<211> 75

<212> PRT

<213> Conus stercusmuscarum

<400> 201

Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe Pro  
 1 5 10 15

Leu Thr Ala Leu Pro Met Asp Gly Asp Gln Pro Ala Asp Gln Pro Ala  
 20 25 30

Asp Arg Met Gln Asp Asp Ile Ser Ser Glu Gln Tyr Pro Leu Phe Asp  
 35 40 45

Lys Arg Gln Lys Cys Cys Thr Gly Lys Lys Gly Ser Cys Ser Gly Lys  
 50 55 60

Ala Cys Lys Asn Leu Lys Cys Cys Ser Gly Arg  
 65 70 75

<210> 202

<211> 23

<212> PRT

<213> Conus stercusmuscarum

<220>

<221> PEPTIDE

<222> (1)..(23)

<223> Xaa at residue 1 is Gln or pyro-Glu

<400> 202

Xaa Lys Cys Cys Thr Gly Lys Lys Gly Ser Cys Ser Gly Lys Ala Cys  
 1 5 10 15

Lys Asn Leu Lys Cys Cys Ser  
 20

<210> 203

<211> 316

<212> DNA

<213> Conus stercusmuscarum

<400> 203  
 gatcgatagc agttcgtat gtctaaactg ggagtcttgc tgaccatctg tctgcttcgt 60  
 ttccccctta ctgcctttcc gatggatgga gatcaacctg cagaccaacc tgcagatcgt 120  
 atgcagaacg acatttcattc tgagcagtat cccttggatataa 180  
 gccccggcg cgtcatgccc cagatattc aaagacaatt ttatttggatgtt 240  
 atgacaacgt gtcgatgacc aacttcgtta tcacgacttc gccaagtgtc taatgaataa 300  
 gtaaaacgat tgcagt 316  
 <210> 204  
 <211> 73  
 <212> PRT  
 <213> *Conus stercusmuscarum*
  
 <400> 204  
 Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe Pro  
 1 5 10 15  
 Leu Thr Ala Leu Pro Met Asp Gly Asp Gln Pro Ala Asp Gln Pro Ala  
 20 25 30  
 Asp Arg Met Gln Asn Asp Ile Ser Ser Glu Gln Tyr Pro Leu Phe Asp  
 35 40 45  
 Lys Arg Gln Lys Cys Cys Gly Pro Gly Ala Ser Cys Pro Arg Tyr Phe  
 50 55 60  
 Lys Asp Asn Phe Ile Cys Gly Cys Cys  
 65 70  
 <210> 205  
 <211> 23  
 <212> PRT  
 <213> *Conus stercusmuscarum*
  
 <220>  
 <221> PEPTIDE  
 <222> (1)..(23)  
 <223> Xaa at residue 1 is Gln or pyro-Glu; Xaa at residue 6 and 11 is Pyro or Hyp; Xaa at residue 13 is Tyr, 125I-Tyr, mono-iodo-Tyr, di-iodo-Tyr, O-sulpho-Tyr or O-phospho-Ty

<400> 205  
 Xaa Lys Cys Cys Gly Xaa Gly Ala Ser Cys Xaa Arg Xaa Phe Lys Asp  
 1 5 10 15

Asn Phe Ile Cys Gly Cys Cys  
 20

<210> 206  
 <211> 331  
 <212> DNA  
 <213> *Conus striatus*

<400> 206  
 cgacccttca agagggatcg atagcagttc gcgatgtcta aactgggggt attgttgacc 60  
 atctgtctgc ttctgtttcc cttactgtcttcc 120  
 caacttgaag atcgtatgca ggacgacatt tcatctgagc agtatccctc gtttggtagg 180  
 agacaaaaagt gttgcggcga aggctcgtca tgccccaat atttcaaaaa caattttatt 240

tgtggttgtt gttaaatgac aacgtgtcga tgaccaactt cgttatcacy actacgcca 300  
 gtgtcttgc taatgataat aaaatgattc c 331  
 <210> 207  
 <211> 73  
 <212> PRT  
 <213> Conus striatus  
 <400> 207  
 Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe Pro  
 1 5 10 15  
 Leu Thr Ala Leu Pro Met Asp Glu Asp Gln Pro Ala Asp Gln Leu Glu  
 20 25 30  
 Asp Arg Met Gln Asp Asp Ile Ser Ser Glu Gln Tyr Pro Ser Phe Val  
 35 40 45  
 Arg Arg Gln Lys Cys Cys Gly Glu Gly Ser Ser Cys Pro Lys Tyr Phe  
 50 55 60  
 Lys Asn Asn Phe Ile Cys Gly Cys Cys  
 65 70  
 <210> 208  
 <211> 23  
 <212> PRT  
 <213> Conus striatus  
 <220>  
 <221> PEPTIDE  
 <222> (1)..(23)  
 <223> Xaa at residue 1 is Gln or pyro-Glu; Xaa at residue 6 is Glu or gamma-carboxy Glu; Xaa at residue 11 is Pro or Hyp; Xaa at residue 13 is Tyr, 125I-Tyr, mono-iodo-Tyr, di-iodo-Tyr, O-sulpho-Tyr or O-phospho-Ty  
 <400> 208  
 Xaa Lys Cys Cys Gly Xaa Gly Ser Ser Cys Xaa Lys Xaa Phe Lys Asn  
 1 5 10 15  
 Asn Phe Ile Cys Gly Cys Cys  
 20  
 <210> 209  
 <211> 256  
 <212> DNA  
 <213> Conus striatus  
 <400> 209  
 ggatccatga tgtctaaact gggagtcttg ttgaccgtct gtctgcttct gtttccctt 60  
 actgtcttcc cgctggatgg agatcaacct gcagaccgac ctgcagagcg tatgcaggac 120  
 gacatttcat ctgacgagca tcccttgttt gataagagac aaaactgttg caatggggga 180  
 tgctccagca aatggtgca agatcacgca cgttggcgc gtcgatgata acgtgttgat 240  
 gaccaacttt ctgcag 256  
 <210> 210  
 <211> 75  
 <212> PRT

&lt;213&gt; Conus striatus

<400> 210  
 Gly Ser Met Met Ser Lys Leu Gly Val Leu Leu Thr Val Cys Leu Leu  
 1 5 10 15  
 Leu Phe Pro Leu Thr Ala Leu Pro Leu Asp Gly Asp Gln Pro Ala Asp  
 20 25 30  
 Arg Pro Ala Glu Arg Met Gln Asp Asp Ile Ser Ser Asp Glu His Pro  
 35 40 45  
 Leu Phe Asp Lys Arg Gln Asn Cys Cys Asn Gly Gly Cys Ser Ser Lys  
 50 55 60  
 Trp Cys Arg Asp His Ala Arg Cys Cys Gly Arg  
 65 70 75

&lt;210&gt; 211

&lt;211&gt; 20

&lt;212&gt; PRT

&lt;213&gt; Conus striatus

<220>  
 <221> PEPTIDE  
 <222> (1)..(20)  
 <223> Xaa at residue 1 is Gln or pyro-Glu; Xaa at residue 12 is Trp or  
 bromo-Tr

<400> 211  
 Xaa Asn Cys Cys Asn Gly Gly Cys Ser Ser Lys Xaa Cys Arg Asp His  
 1 5 10 15

Ala Arg Cys Cys  
 20

<210> 212  
 <211> 235  
 <212> DNA  
 <213> Conus tessulatus

<400> 212  
 ggatccatga tgtctaaact gggagtcttg ttgaccatgt gtctgcttct gtttcccctt 60  
 actgctgttc cgctggatgg agatcaacct gcagaccgac ctgcagagcg taggcaggac 120  
 attgcaactg acgatcatcc tttgtttgat cccgtcaaac ggtgctgcca caaatgctat 180  
 atgggatgca tccctgttg catttagtaa cgtgttgatg accaactttc tcgag 235

<210> 213  
 <211> 68  
 <212> PRT  
 <213> Conus tessulatus

<400> 213  
 Gly Ser Met Met Ser Lys Leu Gly Val Leu Leu Thr Met Cys Leu Leu  
 1 5 10 15

Leu Phe Pro Leu Thr Ala Val Pro Leu Asp Gly Asp Gln Pro Ala Asp  
 20 25 30

Arg Pro Ala Glu Arg Arg Gln Asp Ile Ala Thr Asp Asp His Pro Leu  
 35 40 45

Phe Asp Pro Val Lys Arg Cys Cys His Lys Cys Tyr Met Gly Cys Ile  
 50 55 60

Pro Cys Cys Ile  
 65

<210> 214  
 <211> 14  
 <212> PRT  
 <213> Conus tessulatus

<220>  
 <221> PEPTIDE  
 <222> (1)..(14)  
 <223> Xaa at residue 11 is Pro or Hyp; Xaa at residue 6 is Tyr, 125I-Ty  
 r, mono-iodo-Tyr, di-iodo-Tyr, O-sulpho-Tyr or O-phospho-Ty

<400> 214  
 Cys Cys His Lys Cys Xaa Met Gly Cys Ile Xaa Cys Cys Ile  
 1 5 10

<210> 215  
 <211> 238  
 <212> DNA  
 <213> Conus tessulatus

<400> 215  
 ggatccatga tgtctaaact gggagtcttg ttgaccatct gtgtgcttct gtttcccctt 60  
 actgctgttc cgctggatgg agatcaacct gcagaccaac ctgcagagcg tacgcagaac 120  
 gagcagcatc ccttgtatga tcagaaaaga aagtgttgcc ggccgcccattg cgccatgagc 180  
 tgcggcatgg ctaggtgttg ctattaatga taacgtgttg atgaccaact ttctcgag 238

<210> 216  
 <211> 68  
 <212> PRT  
 <213> Conus tessulatus

<400> 216  
 Gly Ser Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Val Leu  
 1 5 10 15

Leu Phe Pro Leu Thr Ala Val Pro Leu Asp Gly Asp Gln Pro Ala Asp  
 20 25 30

Gln Pro Ala Glu Arg Thr Gln Asn Glu Gln His Pro Leu Tyr Asp Gln  
 35 40 45

Lys Arg Lys Cys Cys Arg Pro Pro Cys Ala Met Ser Cys Gly Met Ala  
 50 55 60

Arg Cys Cys Tyr  
 65

<210> 217  
 <211> 18  
 <212> PRT  
 <213> Conus tessulatus

<220>  
 <221> PEPTIDE  
 <222> (1)..(18)  
 <223> Xaa at residue 5 and 6 is Pro or Hyp; Xaa at residue 18 is Tyr, 1

25I-Tyr, mono-iodo-Tyr, di-iodo-Tyr, O-sulpho-Tyr or O-phospho-Ty

Cys Xaa

<210> 218  
<211> 564  
<212> DNA  
<213> *Conus textile*

<400> 218  
gagtcaaccc actgtcacgc caagagcgg a cggcacagct aaggcaagaa ggatcgatag 60  
cagttcatga tgtctaaact gggagccttg ttgaccatct gtctacttct gtttccctt 120  
actgctgttc cgctggatgg agatcaacat gcagaccaac ctgcacagcg tctgcaggac 180  
cgcattccaa ctgaagatca tcccttattt gatcccaaca aacggtgttgc cccgcccgtg 240  
gcatgcaaca tggatgcaa gccttggatgtt ggttgcaccag ctttgcattc gcggctctat 300  
gaagtgtctta atgaataagt aaaacgattt cagttcggtt cagatttgcgtt gttgtat 360  
ggtctaaaga ttaatgacca aactgttctt ttgatccggaa ttttcacgtt ttttcgtt 420  
cctattcaac actagataag ttaatcacgaa cagatctgtt tttccatcaa tgccttgcgtt 480  
tttggtctgtt catataaattt ttgtttat ttaatttctc gtcaatttca acacgcacac 540  
acacacacac acacacacacac 564

<210> 219  
<211> 69  
<212> PRT  
<213> *Conus textile*

<400> 219  
Met Met Ser Lys Leu Gly Ala Leu Leu Thr Ile Cys Leu Leu Leu Phe  
1 5 10 15

Ser Leu Thr Ala Val Pro Leu Asp Gly Asp Gln His Ala Asp Gln Pro  
20 25 30

Ala Gln Arg Leu Gln Asp Arg Ile Pro Thr Glu Asp His Pro Leu Phe  
35 40 45

Asp Pro Asn Lys Arg Cys Cys Pro Pro Val Ala Cys Asn Met Gly Cys  
50 55 60

Lys Pro Cys Cys Gly  
65

<210> 220  
<211> 16  
<212> PRT  
<213> *Conus textile*

<220>  
<221> PEPTIDE  
<222> (1)..(16)  
<223> Xaa at residue 3, 4 and 13 is Pro or Hyp

<400> 220  
Cys Cys Xaa Xaa Val Ala Cys Asn Met Gly Cys Lys Xaa Cys Cys Gly  
1 5 10 15

<210> 221  
<211> 780  
<212> DNA  
<213> *Conus textile*

<400> 221  
ggatccagac gacaagaag agtcaaccca ctgccacgtc aagagcagag cccacagcta 60  
agacaagaag gatcgatagc agttcatgtat gtttaaactg ggagtcttgt tgaccatctg 120  
tctccttctg tttccctta atgctgttcc gttggatgga gatcaacctg cagaccaacc 180  
tgcagagcgt ctgctggacg acatttcatt taaaataat ccctttatg atcccgccaa 240  
acggtgttgc aggacttgcg tcgggtgcac accttggatggttggatgaccag cctcatcaag 300  
tgtctaaacgataaag cgattgcagt ctgcgttgcata tttacttttgcatttctggtc 360  
taaagattaa tgacccaaact ctcttttgcata tccggatgta cttatatttc tcgatttcata 420  
tccaaacgcta gataagctaa tcacgcacaga tctgattttc tgtcaatgcc ttgttttttg 480  
gtctctcata tcactcttgcatttgcata tttctcgatca ctatataat atatacacac 540  
acacacacac ggaattccgat ttttccagta ccgttcttgg gatcgaggta ttgtgcgtat 600  
ggcttattct gtactctttt ctctcgatca tgatagtgat gtcttctact cccatctgtg 660  
ctacccctgg cttgtatctttt gataggcgat tggcccttac tggttataaaa cccctctgtat 720  
cctactctctt qgacqccctcq qggggcccaac ctccaaataa aqcgacatcc aatgaaaaaa 780

<210> 222  
<211> 66  
<212> PRT  
<213> *Conus textile*

<400> 222  
Met Met Phe Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
1 5 10 15

Ser Leu Asn Ala Val Pro Leu Asp Gly Asp Gln Pro Ala Asp Gln Pro  
20 25 30

Asp Pro Ala Lys Arg Cys Cys Arg Thr Cys Phe Gly Cys Thr Pro Cys  
50 55 60

Cys Gly  
65

<210> 223  
<211> 12  
<212> PRT  
<213> *Conus textile*

<220>  
<221> PEPTIDE

<222> (1)..(12)  
 <223> Xaa at residue 10 is Pro or Hyp  
  
 <400> 223  
 Cys Cys Arg Thr Cys Phe Gly Cys Thr Xaa Cys Cys  
 1 5 10

<210> 224  
 <211> 456  
 <212> DNA  
 <213> Conus textile

<400> 224  
 ggaacagtca accccacagc cacgccaaga gcagacagcc acagctacgt gaagaagggt 60  
 ggagagagggt tcatgtatgtt gaaaatggga gtggtgctat tcatcttct ggtactgttt 120  
 cccctggcaa cgctccagct ggatgcagat caacctgttag aacgatatgc ggagaacaaa 180  
 cagctcctca acccagatga aaggagggaa atcctattgc ctgctctgag gaagttctgc 240  
 tgtgattcga attggtgcca catttcggat tgtgagtgtct gctacggta gcgcgaaca 300  
 tccatggcac tgtgctgggc ggtttcatcc caacaacgac agcgtttgtt gatttcatgt 360  
 atcattgcgc ccacgtctct tgtctaagaa tgacgaacat gattgcactc tggttcagat 420  
 ttctgtttct tttctgacaa taaatgacaa acctcc 456

<210> 225  
 <211> 70  
 <212> PRT  
 <213> Conus textile

<400> 225  
 Met Met Leu Lys Met Gly Val Val Leu Phe Ile Phe Leu Val Leu Phe  
 1 5 10 15

Pro Leu Ala Thr Leu Gln Leu Asp Ala Asp Gln Pro Val Glu Arg Tyr  
 20 25 30

Ala Glu Asn Lys Gln Leu Leu Asn Pro Asp Glu Arg Arg Glu Ile Leu  
 35 40 45

Leu Pro Ala Leu Arg Lys Phe Cys Cys Asp Ser Asn Trp Cys His Asp  
 50 55 60

Cys Glu Cys Cys Tyr Gly  
 65 70

<210> 226  
 <211> 17  
 <212> PRT  
 <213> Conus textile

<220>  
 <221> PEPTIDE  
 <222> (1)..(17)  
 <223> Xaa at residue 14 is Glu or gamma-carboxy Glu; Xaa at residue 7 is Trp or bromo-Trp; Xaa at residue 17 is Tyr, 125I-Tyr, mono-iodo-Tyr, di-iodo-Tyr, O-sulpho-Tyr or O-phospho-Tyr

<400> 226  
 Phe Cys Cys Asp Ser Asn Xaa Cys His Ile Ser Asp Cys Xaa Cys Cys  
 1 5 10 15

Xaa

<210> 227  
 <211> 456  
 <212> DNA  
 <213> Conus textile

<220>  
 <221> misc\_feature  
 <222> (1)..(456)  
 <223> n may be any nucleotide

<400> 227  
 caaggaacag tcaacccac agccacgcca agagcaagaca gccacagcta cgtgaagaag 60  
 ggtggagaga gttcgtat gttaaaaatg ggagtggtgc tattcatctt cctggtactg 120  
 tttccctgg caacgctcca gctggatgca gatcaacctg tagaacgata tgcggagaac 180  
 aaacagctcc tcagcccaga tgaaaggagg gaaatcatat tgcattgtct ggggacgcga 240  
 tgctgttctt gggatgtgtg cgaccacccg agttgtactt gctgcggta ggcggaaaca 300  
 tccatggcgc tgcgtgggc ggttttatcc caacaacgac agcgttgtt gatttcatgt 360  
 atcattgcgc ccacgtctct tgcgttaagaa tgacgaacat gattgcactc tggttcagat 420  
 ttcgtgttct tttctgacaa taaatgacaa aacncc 456

<210> 228  
 <211> 70  
 <212> PRT  
 <213> Conus textile

<400> 228  
 Met Leu Lys Met Gly Val Val Leu Phe Ile Phe Leu Val Leu Phe Pro  
 1 5 10 15

Leu Ala Thr Leu Gln Leu Asp Ala Asp Gln Pro Val Glu Arg Tyr Ala  
 20 25 30

Glu Asn Lys Gln Leu Leu Ser Pro Asp Glu Arg Arg Glu Ile Ile Leu  
 35 40 45

His Ala Leu Gly Thr Arg Cys Cys Ser Trp Asp Val Cys Asp His Pro  
 50 55 60

Ser Cys Thr Cys Cys Gly  
 65 70

<210> 229  
 <211> 15  
 <212> PRT  
 <213> Conus textile

<220>  
 <221> PEPTIDE  
 <222> (1)..(15)  
 <223> Xaa at residue 10 is Pro or Hyp; Xaa at residue 4 is Trp or bromo  
 -Tr.

<400> 229  
 Cys Cys Ser Xaa Asp Val Cys Asp His Xaa Ser Cys Thr Cys Cys  
 1 5 10 15

<210> 230  
 <211> 235  
 <212> DNA  
 <213> Conus textile

<400> 230  
 ggatccatga tgcctaaact gggagtcttg ttgaccatct gtctgcttct gtttcccctt 60  
 actgccttc cgctggatgg agatcaaccc gcagaccaag ctgcagagcg tatgcaggcc 120  
 gagcagcatc ccttgttga tcagaaaaga cggtgctgca agttccatg ccccgatagt 180  
 tgcagatatt tgcgttgccg gtgatgataa cgtgttgatg accaacttcc tcgag 235

<210> 231  
 <211> 67  
 <212> PRT  
 <213> Conus textile

<400> 231  
 Gly Ser Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu  
 1 5 10 15

Leu Phe Pro Leu Thr Ala Leu Pro Leu Asp Gly Asp Gln Pro Ala Asp  
 20 25 30

Gln Ala Ala Glu Arg Met Gln Ala Glu Gln His Pro Leu Phe Asp Gln  
 35 40 45

Lys Arg Arg Cys Cys Lys Phe Pro Cys Pro Asp Ser Cys Arg Tyr Leu  
 50 55 60

Cys Cys Gly  
 65

<210> 232  
 <211> 16  
 <212> PRT  
 <213> Conus textile

<220>  
 <221> PEPTIDE  
 <222> (1)..(16)  
 <223> Xaa at residue 3 and 8 is Pro or Hyp; Xaa at residue 13 is Tyr, 1  
 25I-Tyr, mono-iodo-Tyr, di-iodo-Tyr, O-sulpho-Tyr or O-phospho-Ty

<400> 232  
 Arg Cys Cys Lys Phe Xaa Cys Xaa Asp Ser Cys Arg Xaa Leu Cys Cys  
 1 5 10 15

<210> 233  
 <211> 321  
 <212> DNA  
 <213> Conus tulipa

<400> 233  
 cgacctcaag agggatcgat agcagttcat gtctaaactg ggagtcttgt tgacaatctg 60  
 tctgcttctg tttccctta ctgctctgcc gatggatgg aatggacccctg cagaccgacc 120  
 tgcagagcgt atgcaggaca acatttcata tgagcagcat cccttgggttgg aggagagaca 180  
 cggatgttgc aaggggcccg aaggatgctc ctccagagaa tgcagacccc aacattgttgc 240

cggtcgacga taacgtgtt agggccaaact ttgttatcac ggctacgtca agtgttttagt 300  
 gaataagtaa aatgattgca g 321  
 <210> 234  
 <211> 74  
 <212> PRT  
 <213> *Conus tulipa*  
 <400> 234  
 Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe Pro 1  
 5 10 15  
 Leu Thr Ala Leu Pro Met Asp Gly Asp Glu Pro Ala Asp Arg Pro Ala 20  
 25 30  
 Glu Arg Met Gln Asp Asn Ile Ser Ser Glu Gln His Pro Leu Phe Glu 35  
 40 45  
 Glu Arg His Gly Cys Cys Lys Gly Pro Glu Gly Cys Ser Ser Arg Glu 50  
 55 60  
 Cys Arg Pro Gln His Cys Cys Gly Arg Arg 65  
 70  
 <210> 235  
 <211> 21  
 <212> PRT  
 <213> *Conus tulipa*  
 <220>  
 <221> PEPTIDE  
 <222> (1)..(21)  
 <223> Xaa at residue 8 and 14 is Glu or gamma-carboxy Glu; Xaa at residue 7 and 17 is Pro or Hy  
 <400> 235  
 His Gly Cys Cys Lys Gly Xaa Xaa Gly Cys Ser Ser Arg Xaa Cys Arg 1  
 5 10 15  
 Xaa Gln His Cys Cys 20  
 <210> 236  
 <211> 287  
 <212> DNA  
 <213> *Conus figulinus*  
 <400> 236  
 caagaaggat cgatagcagt tcatgtatgtc taaaactggga gtcttgctga ccatctgtct 60  
 gcttctgatt ccccttactg ctctttcgct ggatggagat caacctgcag accgacacctgc 120  
 agagcgtatg caggatggaa tttcatctga acagcatccc atgtttgatec ccgtcagacg 180  
 gtgttgcccg tggccatgca acataggatg cgtaccttgt tggatgac cagttttgtt 240  
 atcgcggcct catcaaatgt ctaatgaata agtaaaacga ttgcagt 287  
 <210> 237  
 <211> 67  
 <212> PRT  
 <213> *Conus figulinus*  
 <400> 237

Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Ile  
 1 5 10 15

Pro Leu Thr Ala Leu Ser Leu Asp Gly Asp Gln Pro Ala Asp Arg Pro  
 20 25 30

Ala Glu Arg Met Gln Asp Gly Ile Ser Ser Glu Gln His Pro Met Phe  
 35 40 45

Asp Pro Val Arg Arg Cys Cys Pro Trp Pro Cys Asn Ile Gly Cys Val  
 50 55 60

Pro Cys Cys  
 65

<210> 238

<211> 14

<212> PRT

<213> Conus figulinus

<220>

<221> PEPTIDE

<222> (1)..(14)

<223> Xaa at residue 3, 5 and 12 is Pro or Hyp; Xaa at residue 4 is Trp  
 or bromo-Tr

<400> 238

Cys Cys Xaa Xaa Xaa Cys Asn Ile Gly Cys Val Xaa Cys Cys  
 1 5 10

<210> 239

<211> 283

<212> DNA

<213> Conus figulinus

<400> 239

caagaggat cgatagcagt tcatgatgtt taaactggga gtcctgttga ccatctgtat 60  
 gcttctgttt ccctttactg ctcttccgct ggatggagag caacctgcag accaacctgc 120  
 agagcgcatg cagtagatgaca tggtagtgc aatgaatccc tggtttgc tccgtcaaaag 180  
 gtgctgctcg aagaactgctcg cagtagatgcat cccttgc tccgtcaactga ccagcttgc 240  
 tatcgccggcc aaggctctaa tgaataagta aaacgattgc agt 283

<210> 240

<211> 67

<212> PRT

<213> Conus figulinus

<400> 240

Met Met Phe Lys Leu Gly Val Leu Leu Thr Ile Cys Met Leu Leu Phe  
 1 5 10 15

Pro Phe Thr Ala Leu Pro Leu Asp Gly Glu Gln Pro Ala Asp Gln Pro  
 20 25 30

Ala Glu Arg Met Gln Tyr Asp Met Leu Arg Ala Met Asn Pro Trp Phe  
 35 40 45

Asp Pro Val Lys Arg Cys Cys Ser Lys Asn Cys Ala Val Cys Ile Pro  
 50 55 60

Cys Cys Pro

65

<210> 241  
 <211> 14  
 <212> PRT  
 <213> Conus figulinus

<220>  
 <221> PEPTIDE  
 <222> (1)..(14)  
 <223> Xaa at residue 11 and 14 is Pro or Hyp

<400> 241  
 Cys Cys Ser Lys Asn Cys Ala Val Cys Ile Xaa Cys Cys Xaa  
 1 5 10

<210> 242  
 <211> 286  
 <212> DNA  
 <213> Conus figulinus

<400> 242  
 caagaggat cgatagcagt tcatgatgtc taaaactgaga gtcttgttga ccttatgtct 60  
 gcttctgtt ccccttactg ctcttccgct gaatgaagat caacctgcag agcgtatgca 120  
 ggacgacaat tcatctgagc agcaccctt gtatgaccac aaacgaaagt gttgccggtg 180  
 gccatgcccc gcaagatgcg gctttgttg cctgtaataa cgtgtggcc aactttgtta 240  
 tcacggccac gtcaaatgtt taatgaataa gtaaaacgat tgcagt 286

<210> 243  
 <211> 64  
 <212> PRT  
 <213> Conus figulinus

<400> 243  
 Met Met Ser Lys Leu Arg Val Leu Leu Thr Leu Cys Leu Leu Phe  
 1 5 10 15

Pro Leu Thr Ala Leu Pro Leu Asn Glu Asp Gln Pro Ala Glu Arg Met  
 20 25 30

Gln Asp Asp Asn Ser Ser Glu Gln His Pro Leu Tyr Asp His Lys Arg  
 35 40 45

Lys Cys Cys Arg Trp Pro Cys Pro Ala Arg Cys Gly Ser Cys Cys Leu  
 50 55 60

<210> 244  
 <211> 15  
 <212> PRT  
 <213> Conus figulinus

<220>  
 <221> PEPTIDE  
 <222> (1)..(15)  
 <223> Xaa at residue 5 and 7 is Pro or Hyp; Xaa at residue 4 is Trp or  
 bromo-Tr

<400> 244  
 Cys Cys Arg Xaa Xaa Cys Xaa Ala Arg Cys Gly Ser Cys Cys Leu  
 1 5 10 15

<210> 245

&lt;211&gt; 301

&lt;212&gt; DNA

&lt;213&gt; Conus figulinus

&lt;400&gt; 245

caagaggat cgatagcagt tcatgatgtc taaaactggga gtcttggta ccttatgtct 60  
 gcttctgttt cccctgactg ctcttccgct gcatgaatg caagctgcag accgacccgc 120  
 agagcgtatg cagggcatgt catctgaaca gcattccctc tttgatcccg tcaaaccgg 180  
 ttgcgagttg tcacgctgcc ttggatgcgt cccttggta acatcttaat aacgtgtg 240  
 tgaccaactg tgttatcacg gccacgtcaa gtgtctaatg aataagtaaa atgattgcag 300  
 t 301

&lt;210&gt; 246

&lt;211&gt; 68

&lt;212&gt; PRT

&lt;213&gt; Conus figulinus

&lt;400&gt; 246

Met	Met	Ser	Lys	Leu	Gly	Val	Leu	Leu	Thr	Leu	Cys	Leu	Leu	Phe	
1				5				10			15				
Pro	Leu	Thr	Ala	Leu	Pro	Leu	Asp	Glu	Asp	Gln	Ala	Ala	Asp	Arg	Pro
					20			25			30				
Ala	Glu	Arg	Met	Gln	Gly	Met	Ser	Ser	Glu	Gln	His	Pro	Phe	Phe	Asp
			35				40				45				
Pro	Val	Lys	Arg	Cys	Cys	Glu	Leu	Ser	Arg	Cys	Leu	Gly	Cys	Val	Pro
					50		55			60					

Cys Cys Thr Ser

65

&lt;210&gt; 247

&lt;211&gt; 16

&lt;212&gt; PRT

&lt;213&gt; Conus figulinus

&lt;220&gt;

&lt;221&gt; PEPTIDE

&lt;222&gt; (1)..(16)

&lt;223&gt; Xaa at residue 3 and 12 is Pro or Hyp

&lt;400&gt; 247

Cys	Cys	Xaa	Leu	Ser	Arg	Cys	Leu	Gly	Cys	Val	Xaa	Cys	Cys	Thr	Ser
1				5				10			15				

&lt;210&gt; 248

&lt;211&gt; 301

&lt;212&gt; DNA

&lt;213&gt; Conus figulinus

&lt;400&gt; 248

caagaggat cgatagcagt tcatgatgtc taaaactggga gtcttggta ccttatgtct 60  
 gcttctgttt cccctgactg ctcttccgct gcatgaatg caacctgcag accgacccgc 120  
 agagcgtatg cagggcatgt catctgaaca gcattccctc tttgatcccg tcaaaccgg 180  
 ttgcgagttg tcaaaatgcc atggatgcgt cccttggta acatcttaat aacgtgcgg 240

tgaccaactg tgttatcacg gccacgtcaa gtgtctaatt aataagtaaa atgattgcag 300  
 t 301  
 <210> 249  
 <211> 68  
 <212> PRT  
 <213> Conus figulinus  
 <400> 249  
 Met Met Ser Lys Leu Gly Val Leu Leu Thr Leu Cys Leu Leu Leu Phe  
 1 5 10 15  
 Pro Leu Thr Ala Leu Pro Leu Asp Glu Asp Gln Pro Ala Asp Arg Pro  
 20 25 30  
 Ala Glu Arg Met Gln Gly Met Ser Ser Glu Gln His Pro Phe Phe Asp  
 35 40 45  
 Pro Val Lys Arg Cys Cys Glu Leu Ser Lys Cys His Gly Cys Val Pro  
 50 55 60  
 Cys Cys Ile Pro  
 65  
 <210> 250  
 <211> 16  
 <212> PRT  
 <213> Conus figulinus  
 <220>  
 <221> PEPTIDE  
 <222> (1)..(16)  
 <223> Xaa at residue 3 is Glu or gamma-carboxy Glu; Xaa at residue 12 and 16 is Pro or Hy  
 <400> 250  
 Cys Cys Xaa Leu Ser Lys Cys His Gly Cys Val Xaa Cys Cys Ile Xaa  
 1 5 10 15  
 <210> 251  
 <211> 298  
 <212> DNA  
 <213> Conus quercinus  
 <400> 251  
 caagagggtat ctagatcgt taaactcgga gtcttggta ccatctgtct 60  
 gtttctgttt ccccttacag ctcttcagct ggatggagat caacctgcag accgacactgc 120  
 agagcgtacg caggacattt catctgaaca gtatcgaaag tttgatcaga gacagaggta 180  
 ttggccgtgg ccatgccccg gtagttgcag atgctccgt tatcgtaac gtgtggta 240  
 ccagctttgt tatcacgacc acgccaagtg tctaacaat aagtaaaatg attgcagt 298  
 <210> 252  
 <211> 68  
 <212> PRT  
 <213> Conus quercinus  
 <400> 252  
 Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Val Leu Phe  
 1 5 10 15  
 Pro Leu Thr Ala Leu Gln Leu Asp Gly Asp Gln Pro Ala Asp Arg Pro

73

20

25

30

Ala Glu Arg Thr Gln Asp Ile Ser Ser Glu Gln Tyr Arg Lys Phe Asp  
 35 40 45

Gln Arg Gln Arg Cys Cys Arg Trp Pro Cys Pro Gly Ser Cys Arg Cys  
 50 55 60

Cys Arg Tyr Arg  
 65

<210> 253  
 <211> 18  
 <212> PRT  
 <213> Conus quercinus

<220>  
 <221> PEPTIDE  
 <222> (1)..(18)  
 <223> Xaa at residue 1 is Gln or pyro-Glu; Xaa at residue 7 and 9 is Pr o or Hyp; Xaa at residue 6 is Trp or bromo-Trp; Xaa at residue 17 is Tyr, 125I-Tyr, mono-iodo-Tyr, di-iodo-Tyr, O-sulpho-Tyr or O-phospho-Ty

<400> 253  
 Xaa Arg Cys Cys Arg Xaa Xaa Cys Xaa Gly Ser Cys Arg Cys Cys Arg  
 1 5 10 15

Xaa Arg

<210> 254  
 <211> 313  
 <212> DNA  
 <213> Conus quercinus

<400> 254  
 caagaggat cgatagcagt tcatgatgtc taaaactggga gtcttggta ccatctgtct 60  
 gcttctgttt ccccttactg ctcttccact ggtggagat caacctgcag atcaatctgc 120  
 agagcgacct gcagagcgta cgcaggacgca cattcagcag catccgttat atgatccgaa 180  
 aagaagggtg tgccgttatac catgccccga cagctccac ggatcttgct gctataagt 240  
 ataacatgtt gatggccagc tttgttatca cggccacgta aagtgtctaa tgaataagta 300  
 aaacgattgc agt 313

<210> 255  
 <211> 72  
 <212> PRT  
 <213> Conus quercinus

<400> 255  
 Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Phe  
 1 5 10 15

Pro Leu Thr Ala Leu Pro Leu Asp Gly Asp Gln Pro Ala Asp Gln Ser  
 20 25 30

Ala Glu Arg Pro Ala Glu Arg Thr Gln Asp Asp Ile Gln Gln His Pro  
 35 40 45

Leu Tyr Asp Pro Lys Arg Arg Cys Cys Arg Tyr Pro Cys Pro Asp Ser  
 50 55 60

Cys His Gly Ser Cys Cys Tyr Lys  
65 70

<210> 256  
<211> 18  
<212> PRT  
<213> Conus quercinus

<220>  
<221> PEPTIDE  
<222> (1)..(18)  
<223> Xaa at residue 6 and 8 is Pro or Hyp; Xaa at residue 5 and 17 is Tyr, 125I-Tyr, mono-iodo-Tyr, di-iodo-Tyr, O-sulpho-Tyr or O-phospho-Ty

<400> 256  
Arg Cys Cys Arg Xaa Xaa Cys Xaa Asp Ser Cys His Gly Ser Cys Cys  
1 5 10 15

Xaa Lys

<210> 257  
<211> 256  
<212> DNA  
<213> Conus wittigi

<400> 257  
ggatccatga tgtctaaact gggagtcttg ttgaccatct gtctgcttct gtttcccatt 60  
actgctcttc cgggtgggtgg agatcagcct gcagaccgac ttgcagagcg tatgcaggac 120  
gacacttcat ctgagcagca tccctttgaa aagagactac catcatgttg cgactttgag 180  
aggctttgcg tagtaccagc atgcatacgt catcagtgtt gcacaggata acgtgttgat 240  
gaccaacttt ctgcag 256

<210> 258  
<211> 74  
<212> PRT  
<213> Conus wittigi

<400> 258  
Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
1 5 10 15

Pro Ile Thr Ala Leu Pro Val Gly Gly Asp Gln Pro Ala Asp Arg Leu  
20 25 30

Ala Glu Arg Met Gln Asp Asp Thr Ser Ser Glu Gln His Pro Phe Glu  
35 40 45

Lys Arg Leu Pro Ser Cys Cys Asp Phe Glu Arg Leu Cys Val Val Pro  
50 55 60

Ala Cys Ile Arg His Gln Cys Cys Thr Gly  
65 70

<210> 259  
<211> 23  
<212> PRT  
<213> Conus wittigi  
  
<220>

&lt;221&gt; PEPTIDE

&lt;222&gt; (1)..(23)

&lt;223&gt; Xaa at residue 8 is Glu or gamma-carboxy Glu; Xaa at residue 2 and 14 is Pro or Hyp

&lt;400&gt; 259

Leu Xaa Ser Cys Cys Asp Phe Xaa Arg Leu Cys Val Val Xaa Ala Cys  
1 5 10 15Ile Arg His Gln Cys Cys Thr  
20

&lt;210&gt; 260

&lt;211&gt; 14

&lt;212&gt; PRT

&lt;213&gt; Conus betulinus

&lt;220&gt;

&lt;221&gt; PEPTIDE

&lt;222&gt; (1)..(14)

&lt;223&gt; Xaa at residue 11 is Pro or Hyp; Xaa at residue 14 is Trp or bromo-Tr

&lt;400&gt; 260

Cys Cys Lys Gln Ser Cys Thr Thr Cys Met Xaa Cys Cys Xaa  
1 5 10

&lt;210&gt; 261

&lt;211&gt; 259

&lt;212&gt; DNA

&lt;213&gt; Conus tulipa

&lt;220&gt;

&lt;221&gt; misc\_feature

&lt;222&gt; (1)..(259)

&lt;223&gt; n may be any nucleotide

&lt;400&gt; 261

ggatccatga tgtctaaact gggagtcttg ttgacaatct gtctgcttct gtttcccctt 60

actgctctgc cgatggatgg agatgaacct gcagaccgac ctgcagagcg tatgcaggac 120

aacatttcat ctgagcagca tcccttggtt gaggagagac acggatgttg cgaggggccg 180

aaggatgtct cctccagaga atgcagaccc caacattgtt gcggtcgacg ataacgtgtt 240

gatgaccaac tntctcgag 259

&lt;210&gt; 262

&lt;211&gt; 75

&lt;212&gt; PRT

&lt;213&gt; Conus tulipa

&lt;400&gt; 262

Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
1 5 10 15Pro Leu Thr Ala Leu Pro Met Asp Gly Asp Glu Pro Ala Asp Arg Pro  
20 25 30Ala Glu Arg Met Gln Asp Asn Ile Ser Ser Glu Gln His Pro Leu Phe  
35 40 45Glu Glu Arg His Gly Cys Cys Glu Gly Pro Lys Gly Cys Ser Ser Arg  
50 55 60

Glu Cys Arg Pro Gln His Cys Cys Gly Arg Arg  
 65 70 75

<210> 263  
 <211> 21  
 <212> PRT  
 <213> *Conus tulipa*

<220>  
 <221> PEPTIDE  
 <222> (1)..(21)  
 <223> Xaa at residue 5 and 14 is Glu or gamma-carboxy Glu; Xaa at residue 7 and 17 is Pro or Hy

<400> 263  
 His Gly Cys Cys Xaa Gly Xaa Lys Gly Cys Ser Ser Arg Xaa Cys Arg  
 1 5 10 15

Xaa Gln His Cys Cys  
 20

<210> 264  
 <211> 262  
 <212> DNA  
 <213> *Conus aurisiacus*

<220>  
 <221> misc feature  
 <222> (1)..(262)  
 <223> n may be any nucleotide

<400> 264  
 ggatccatga tgtctaaact gggagtcttg ttgaccatct gtctacttct gtttccctt 60  
 actgctttc cgatggatgg agatcaacct gcagaccaac ctgcagatcg tatgcaggac 120  
 gacatttcat ctgagcagta tcccttgttt gataagagac aaaagtgttg cactggagg 180  
 aagggtcat gctccggcaa agcatgcaaa aatctcaa at gttgctctgg acgataacgt 240  
 gttgatgacc aactttctcg an 262

<210> 265  
 <211> 76  
 <212> PRT  
 <213> *Conus aurisiacus*

<400> 265  
 Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15

Pro Leu Thr Ala Phe Pro Met Asp Gly Asp Gln Pro Ala Asp Gln Pro  
 20 25 30

Ala Asp Arg Met Gln Asp Asp Ile Ser Ser Glu Gln Tyr Pro Leu Phe  
 35 40 45

Asp Lys Arg Gln Lys Cys Cys Thr Gly Arg Lys Gly Ser Cys Ser Gly  
 50 55 60

Lys Ala Cys Lys Asn Leu Lys Cys Cys Ser Gly Arg  
 65 70 75

<210> 266

<211> 23  
 <212> PRT  
 <213> Conus aurisiacus

<220>  
 <221> PEPTIDE  
 <222> (1)...(23)  
 <223> Xaa at residue 1 is Gln or pyro-Glu

<400> 266  
 Xaa Lys Cys Cys Thr Gly Arg Lys Gly Ser Cys Ser Gly Lys Ala Cys  
 1 5 10 15  
 Lys Asn Leu Lys Cys Cys Ser  
 20

<210> 267  
 <211> 239  
 <212> DNA  
 <213> Conus betulinus

<400> 267  
 ggatccatga tgtctaaact gggagtcttg ttgaccatct gtctgcttct gtttcccctt 60  
 actgctgttc cgttggatgg agatcaacct gcagaccáac ctgcagagcg tatgcagaac 120  
 gagcagcatc ctcgtttga tcagaaaaga aggtgctgcc ggtggccatg ccccagtata 180  
 tgcggcatgg ctaggtgttg cttcgtcatg ataacgtgtt gatgaccaac tttctcgag 239

<210> 268  
 <211> 71  
 <212> PRT  
 <213> Conus betulinus

<400> 268  
 Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15  
 Pro Leu Thr Ala Val Pro Leu Asp Gly Asp Gln Pro Ala Asp Gln Pro  
 20 25 30

Ala Glu Arg Met Gln Asn Glu Gln His Pro Ser Phe Asp Gln Lys Arg  
 35 40 45

Arg Cys Cys Arg Trp Pro Cys Pro Ser Ile Cys Gly Met Ala Arg Cys  
 50 55 60

Cys Phe Val Met Ile Thr Cys  
 65 70

<210> 269  
 <211> 23  
 <212> PRT  
 <213> Conus betulinus

<220>  
 <221> PEPTIDE  
 <222> (1)...(23)  
 <223> Xaa at residue 6 and 8 is Pro or Hyp; Xaa at residue 5 is Trp or  
 bromo-Tr

<400> 269  
 Arg Cys Cys Arg Xaa Xaa Cys Xaa Ser Ile Cys Gly Met Ala Arg Cys  
 1 5 10 15

Cys Phe Val Met Ile Thr Cys  
20

<210> 270  
<211> 226  
<212> DNA  
<213> Conus betulinus

<220>  
<221> misc\_feature  
<222> (1)..(226)  
<223> n may be any nucleotide

<400> 270  
ggatccatga tgtctaaact gggagtcttg ttgatcatct gtctgcttct gtttcccctt 60  
actgctgttc cgctggatgg agatcagcct gcagagcgta cgcagatcga gcagcatccc 120  
ttgtttgacc agaaaagaag gtgttgcgg tggccatgcc ccagtagatg cggcatggct 180  
agggttgtct tcgtcatgat aacgtgttga tgancgacct ctcnag 226

<210> 271  
<211> 67  
<212> PRT  
<213> Conus betulinus

<400> 271  
Met Met Ser Lys Leu Gly Val Leu Leu Ile Ile Cys Leu Leu Leu Phe  
1 5 10 15

Pro Leu Thr Ala Val Pro Leu Asp Gly Asp Gln Pro Ala Glu Arg Thr  
20 25 30

Gln Ile Glu Gln His Pro Leu Phe Asp Gln Lys Arg Arg Cys Cys Arg  
35 40 45

Trp Pro Cys Pro Ser Arg Cys Gly Met Ala Arg Cys Cys Phe Val Met  
50 55 60

Ile Thr Cys  
65

<210> 272  
<211> 23  
<212> PRT  
<213> Conus betulinus

<220>  
<221> PEPTIDE  
<222> (1)..(23)  
<223> Xaa at residue 6 and 8 is Pro or Hyp; Xaa at residue 5 is Trp or  
bromo-Tr

<400> 272  
Arg Cys Cys Arg Xaa Xaa Cys Xaa Ser Arg Cys Gly Met Ala Arg Cys  
1 5 10 15  
Cys Phe Val Met Ile Thr Cys  
20

<210> 273  
<211> 262  
<212> DNA  
<213> Conus parius

<400> 273  
 ggatccatga tgtctaaact gggagtcttg ttgaccatct gtctgcttct gtttcccctt 60  
 actgctcttc cgatggatgg tcatcaaccc gcagaccgac ttgttagagcg tatgcaggac 120  
 aacatttcat ctgagcagca tcccttcttt gaaaagagaa gagggaggctg ttgcacacct 180  
 ccgaagaaat gcaaagaccc agcctgcaaa cctgcacgtt gctgcggccc aggataacgt 240  
 gttgatgacc aactttctcg cc 262

<210> 274  
 <211> 76  
 <212> PRT  
 <213> Conus parius

<400> 274  
 Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15

Pro Leu Thr Ala Leu Pro Met Asp Gly Asp Gln Pro Ala Asp Arg Leu  
 20 25 30

Val Glu Arg Met Gln Asp Asn Ile Ser Ser Glu Gln His Pro Phe Phe  
 35 40 45

Glu Lys Arg Arg Gly Gly Cys Cys Thr Pro Pro Lys Lys Cys Lys Asp  
 50 55 60

Arg Ala Cys Lys Pro Ala Arg Cys Cys Gly Pro Gly  
 65 70 75

<210> 275  
 <211> 24  
 <212> PRT  
 <213> Conus parius

<220>  
 <221> PEPTIDE  
 <222> (1)..(24)  
 <223> Xaa at residue 7, 8, 18 and 24 is Pro or Hyp

<400> 275  
 Arg Gly Gly Cys Cys Thr Xaa Xaa Lys Lys Cys Lys Asp Arg Ala Cys  
 1 5 10 15

Lys Xaa Ala Arg Cys Cys Gly Xaa  
 20

<210> 276  
 <211> 259  
 <212> DNA  
 <213> Conus parius  
 <400> 276  
 ggatccatga tgtctaaact gggagtcttg ttgaccatct gtctgcttct gtttcccctt 60  
 actgctcttc cgatggatgg tcatcaaccc gcagaccgac ttgttagagcg tatgcaggac 120  
 aacatttcat ctgagcagca tcccttcttt gaaaagagaa gagggaggctg cacacctccg 180  
 aggaaatgca aagaccgagc ctgcaaacct gcacgtgtt gcggcccagg ataacgtgtt 240  
 gatgaccaac tttctcgag 259

<210> 277  
 <211> 75  
 <212> PRT  
 <213> Conus parius

<400> 277  
 Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15

Pro Leu Thr Ala Leu Pro Met Asp Gly Asp Gln Pro Ala Asp Arg Leu  
 20 25 30

Val Glu Arg Met Gln Asp Asn Ile Ser Ser Glu Gln His Pro Phe Phe  
 35 40 45

Glu Lys Arg Arg Gly Cys Cys Thr Pro Pro Arg Lys Cys Lys Asp Arg  
 50 55 60

Ala Cys Lys Pro Ala Arg Cys Cys Gly Pro Gly  
 65 70 75

<210> 278  
 <211> 23  
 <212> PRT  
 <213> Conus parius

<220>  
 <221> PEPTIDE  
 <222> (1)..(23)  
 <223> Xaa at residue 6, 7, 17 and 23 is Pro or Hyp

<400> 278  
 Arg Gly Cys Cys Thr Xaa Xaa Arg Lys Cys Lys Asp Arg Ala Cys Lys  
 1 5 10 15

Xaa Ala Arg Cys Cys Gly Xaa  
 20

<210> 279  
 <211> 241  
 <212> DNA  
 <213> Conus coronatus

<400> 279  
 ggatccatga tgtctaaact gggagtcttg ttgaccatct gtctgcttct gtttccaatt 60  
 actgcccttc cgctggatga agatcaacct gcagaccgac ctgcagagcg tatgcaggac 120  
 attgcaactg aacagcatcc cttgtttat cccgtcaaac ggtgctgcga ttggccatgc 180  
 atccccaggat gcaccccttg ttgcttgcct tgataacgtg ttgatgacca actttctcga 240  
 g 241

<210> 280  
 <211> 68  
 <212> PRT  
 <213> Conus coronatus

<400> 280  
 Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15

Pro Ile Thr Ala Leu Pro Leu Asp Glu Asp Gln Pro Ala Asp Arg Pro  
 20 25 30

Ala Glu Arg Met Gln Asp Ile Ala Thr Glu Gln His Pro Leu Phe Asp  
 35 40 45

Pro Val Lys Arg Cys Cys Asp Trp Pro Cys Ile Pro Gly Cys Thr Pro  
 50 55 60

Cys Cys Leu Pro  
 65

<210> 281  
 <211> 16  
 <212> PRT  
 <213> Conus coronatus

<220>  
 <221> PEPTIDE  
 <222> (1)..(16)  
 <223> Xaa at residue 5, 8, 12 and 16 is Pro or Hyp; Xaa at residue 4 is  
 Trp or bromo-Tr

<400> 281  
 Cys Cys Asp Xaa Xaa Cys Ile Xaa Gly Cys Thr Xaa Cys Cys Leu Xaa  
 1 5 10 15

<210> 282  
 <211> 244  
 <212> DNA  
 <213> Conus musicus

<400> 282  
 ggatccatga tgtctaaact gggagtccctt ttgaccatct gtctgcttct gtttcctctt 60  
 tctgcttcc cgatggatga agatcaactt gcagacctac ctgcagagcg tatgcgggac 120  
 actgcaactg tagatcatcc ctcctatgtat cctgacaaaag cgtgctgcga gcagagctgt 180  
 acaacatgct ttccgtgctg ctagccttga acacagtaac gtgttgatga ccaactttct 240  
 cgag 244

<210> 283  
 <211> 65  
 <212> PRT  
 <213> Conus musicus

<400> 283  
 Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15

Pro Leu Ser Ala Leu Pro Met Asp Glu Asp Gln Leu Ala Asp, Leu Pro  
 20 25 30

Ala Glu Arg Met Arg Asp Thr Ala Thr Val Asp His Pro Ser Tyr Asp  
 35 40 45

Pro Asp Lys Ala Cys Cys Glu Gln Ser Cys Thr Thr Cys Phe Pro Cys  
 50 55 60

Cys  
 65

<210> 284  
 <211> 14  
 <212> PRT

<213> Conus musicus

<220>

<221> PEPTIDE

<222> (1)..(14)

<223> Xaa at residue 4 is Glu or gamma-carboxy Glu; Xaa at residue 12 is Pro or Hy

<400> 284

Ala Cys Cys Xaa Gln Ser Cys Thr Thr Cys Phe Xaa Cys Cys  
1 5 10

<210> 285

<211> 14

<212> PRT

<213> Conus betulinus

<220>

<221> PEPTIDE

<222> (1)..(14)

<223> Xaa at residue 4 is Glu or gamma-carboxy Glu; Xaa at residue 12 is Pro or Hy

<400> 285

Ala Cys Cys Xaa Gln Ser Cys Thr Thr Cys Met Xaa Cys Cys  
1 5 10

<210> 286

<211> 14

<212> PRT

<213> Conus betulinus

<220>

<221> PEPTIDE

<222> (1)..(14)

<223> Xaa at residue 3 is Glu or gamma-carboxy Glu; Xaa at residue 11 is Pro or Hyp; Xaa at residue 14 is Trp or bromo-Tr

<400> 286

Cys Cys Xaa Gln Ser Cys Thr Thr Cys Met Xaa Cys Cys Xaa  
1 5 10

<210> 287

<211> 235

<212> DNA

<213> Conus pennaceus

<400> 287

ggatccatga tgtctaaact gggagtcttg ttgaccatct gtctgcttct gtttccctt 60

actgctcttc cgctggatgg agatcaacct gcataccaaag ctgcagagcg tatgcaggcc 120

gagcatcatc ccttggttga tcagaaaaga cggtgctgca agtttccatg ccccgatagt 180

tgcaaatatt tgtgttgcgg gtgatgataa catgttgatg accaactttc ttgag 235

<210> 288

<211> 65

<212> PRT

<213> Conus pennaceus

<400> 288

Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
1 5 10 15

Pro Leu Thr Ala Leu Pro Leu Asp Gly Asp Gln Pro Ala Tyr Gln Ala  
 20 25 30

Ala Glu Arg Met Gln Ala Glu His His Pro Leu Phe Asp Gln Lys Arg  
 35 40 45

Arg Cys Cys Lys Phe Pro Cys Pro Asp Ser Cys Lys Tyr Leu Cys Cys  
 50 55 60

Gly  
 65

<210> 289  
 <211> 16  
 <212> PRT  
 <213> Conus pennaceus

<220>  
 <221> PEPTIDE  
 <222> (1)..(16)  
 <223> Xaa at residue 6 and 8 is Pro or Hyp; Xaa at residue 13 is Tyr, 1  
 25I-Tyr, mono-iodo-Tyr, di-iodo-Tyr, O-sulpho-Tyr or O-phospho-Ty

<400> 289  
 Arg Cys Cys Lys Phe Xaa Cys Xaa Asp Ser Cys Lys Xaa Leu Cys Cys  
 1 5 10 15

<210> 290  
 <211> 241  
 <212> DNA  
 <213> Conus pulicarius

<400> 290  
 ggatccatga tgtctaaact gggagtcttg ttgaccatct gtctgcttct gtttccctt 60  
 actgctttc cgatggatgg ttagtcaactt gcagaccgac ttgttagagcg tatgcaggac 120  
 aacatttcat ctgagcagca tcccttctt gatcccgtaa aacggtgttg cgtcagctgt 180  
 tacatggat gcatcccttg ttgcttcttag taataacgtg ttgatgacca actttctcga 240  
 g 241

<210> 291  
 <211> 67  
 <212> PRT  
 <213> Conus pulicarius

<400> 291  
 Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15

Pro Leu Thr Ala Leu Pro Met Asp Gly Asp Gln Leu Ala Asp Arg Leu  
 20 25 30

Val Glu Arg Met Gln Asp Asn Ile Ser Ser Glu Gln His Pro Phe Phe  
 35 40 45

Asp Pro Val Lys Arg Cys Cys Val Ser Cys Tyr Met Gly Cys Ile Pro  
 50 55 60

Cys Cys Phe  
 65

<210> 292  
 <211> 14  
 <212> PRT  
 <213> Conus pulicarius

<220>  
 <221> PEPTIDE  
 <222> (1)..(14)  
 <223> Xaa at residue 11 is Pro or Hyp; Xaa at residue 6 is Tyr, 125I-Ty  
 r, mono-iodo-Tyr, di-iodo-Tyr, O-sulpho-Tyr or O-phospho-Ty

<400> 292  
 Cys Cys Val Ser Cys Xaa Met Gly Cys Ile Xaa Cys Cys Phe  
 1 5 10

<210> 293  
 <211> 244  
 <212> DNA  
 <213> Conus pulicarius

<400> 293  
 ggatccatga tgtctaaact gggagtcttg ttgaccgtct gtctgcttct gtgtcccctt 60  
 actgtcttcc actggatga agatcaactt gcagaccgac ctgcagagcg tatgcaggat 120  
 gacacttcag ctgcacagat tttcggttt gatcccgatca aacggtgctg caaattgcta 180  
 tgctactcgg gatgcactcc ttgttgccat atttgataac gtgttgatga ccaactttct 240  
 cgag 244

<210> 294  
 <211> 67  
 <212> PRT  
 <213> Conus pulicarius

<400> 294  
 Met Met Ser Lys Leu Gly Val Leu Leu Thr Val Cys Leu Leu Cys  
 1 5 10 15

Pro Leu Thr Ala Leu Pro Leu Asp Glu Asp Gln Leu Ala Asp Arg Pro  
 20 25 30

Ala Glu Arg Met Gln Asp Asp Thr Ser Ala Ala Gln Ile Phe Gly Phe  
 35 40 45

Asp Pro Val Lys Arg Cys Cys Lys Leu Leu Cys Gly Cys Thr Pro Cys  
 50 55 60

Cys His Ile  
 65

<210> 295  
 <211> 16  
 <212> PRT  
 <213> Conus pulicarius

<220>  
 <221> PEPTIDE  
 <222> (1)..(16)  
 <223> Xaa at residue 12 is Pro or Hyp; Xaa at residue 7 is Tyr, 125I-Ty  
 r, mono-iodo-Tyr, di-iodo-Tyr, O-sulpho-Tyr or O-phospho-Ty

<400> 295  
 Cys Cys Lys Leu Leu Cys Xaa Ser Gly Cys Thr Xaa Cys Cys His Ile

85

1 5 10 15

&lt;210&gt; 296

&lt;211&gt; 259

&lt;212&gt; DNA

<213> *Conus rattus*

&lt;400&gt; 296

```
ggatccatga tgtctaaact gggagtcttg ttgaccatct gtctgcttgt gtttccgctt 60
actgctcttc cgatggatgg tgcataacct gcagaccgac ttgttagagcg tatacaggac 120
aacatttcat ctgagcagca tcccttcttt gaaaagagaa gaggctgttg cgcacccctcg 180
aggaaatgca aagaccgac ctgcaaacct gcacgttgc gcggccagg ataacgtgtt 240
gatgaccaac tttctcgag 259
```

&lt;210&gt; 297

&lt;211&gt; 75

&lt;212&gt; PRT

<213> *Conus rattus*

&lt;400&gt; 297

```
Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Val Phe
1 5 10 15
```

```
Pro Leu Thr Ala Leu Pro Met Asp Gly Asp Gln Pro Ala Asp Arg Leu
20 25 30
```

```
Val Glu Arg Ile Gln Asp Asn Ile Ser Ser Glu Gln His Pro Phe Phe
35 40 45
```

```
Glu Lys Arg Arg Gly Cys Cys Ala Pro Pro Arg Lys Cys Lys Asp Arg
50 55 60
```

```
Ala Cys Lys Pro Ala Arg Cys Cys Gly Pro Gly
65 70 75
```

&lt;210&gt; 298

&lt;211&gt; 23

&lt;212&gt; PRT

<213> *Conus rattus*

&lt;220&gt;

&lt;221&gt; PEPTIDE

&lt;222&gt; (1)...(23)

&lt;223&gt; Xaa at residue 6, 7, 17 and 23 is Pro or Hyp

&lt;400&gt; 298

```
Arg Gly Cys Cys Ala Xaa Xaa Arg Lys Cys Lys Asp Arg Ala Cys Lys
1 5 10 15
```

```
Xaa Ala Arg Cys Cys Gly Xaa
20
```

&lt;210&gt; 299

&lt;211&gt; 262

&lt;212&gt; DNA

<213> *Conus stercusmuscarum*

&lt;400&gt; 299

```
ggatccatga tgtctaaact gggagtcttg ttgacaatct gtctgcttgt gtttccgctt 60
attgctcttc cgatggatgg agatcaacct gcagaccgac ctgcagagcg tatgcaggac 120
```

gacatttcat ctgagaagca tcccttgttt gataagagac aacgggtttt caatggcgg 180  
 aggggatgtct ccagcagatg gtgcagagat cactcacgtt gttgcggtcg acgataacgt 240  
 gttgatgacc aactttctcg ag 262

<210> 300  
 <211> 76  
 <212> PRT  
 <213> Conus stercusmuscarum

<400> 300  
 Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15

Pro Leu Ile Ala Leu Pro Leu Asp Gly Asp Gln Pro Ala Asp Arg Pro  
 20 25 30

Ala Glu Arg Met Gln Asp Asp Ile Ser Ser Glu Lys His Pro Leu Phe  
 35 40 45

Asp Lys Arg Gln Arg Cys Cys Asn Gly Arg Arg Gly Cys Ser Ser Arg  
 50 55 60

Trp Cys Arg Asp His Ser Arg Cys Cys Gly Arg Arg  
 65 70 75

<210> 301  
 <211> 22  
 <212> PRT  
 <213> Conus stercusmuscarum

<220>  
 <221> PEPTIDE  
 <222> (1)..(22)  
 <223> Xaa at residue 1 is Gln or pyro-Glu; Xaa at residue 14 is Trp or  
 bromo-Tr

<400> 301  
 Xaa Arg Cys Cys Asn Gly Arg Arg Gly Cys Ser Ser Arg Xaa Cys Arg  
 1 5 10 15

Asp His Ser Arg Cys Cys  
 20

<210> 302  
 <211> 241  
 <212> DNA  
 <213> Conus ebraceus

<400> 302  
 ggatccatga tgtctaaact gggagtcttg ttgaccatct gtctgcttct gtttcccctt 60  
 actgctcttc cactggatga aggtcaacct gcagacctac ctgcagagcg tatgcaggac 120  
 attgcaactg aacagcatcc cttgtttgat cctgtcaaac ggtgttgcga gcagccatgc 180  
 tacatggat gcatcccttg ttgcttctaa taataacgtg ttgatgacca actttctcga 240

g 241

<210> 303  
 <211> 67  
 <212> PRT

&lt;213&gt; Conus ebraceus

&lt;400&gt; 303

Met	Met	Ser	Lys	Leu	Gly	Val	Leu	Leu	Thr	Ile	Cys	Leu	Leu	Phe
1				5				10				15		

Pro	Leu	Thr	Ala	Leu	Pro	Leu	Asp	Glu	Gly	Gln	Pro	Ala	Asp	Leu	Pro
	20				25						30				

Ala	Glu	Arg	Met	Gln	Asp	Ile	Ala	Thr	Glu	Gln	His	Pro	Leu	Phe	Asp
	35				40						45				

Pro	Val	Lys	Arg	Cys	Cys	Glu	Gln	Pro	Cys	Tyr	Met	Gly	Cys	Ile	Pro
	50				55				60						

Cys	Cys	Phe
65		

&lt;210&gt; 304

&lt;211&gt; 15

&lt;212&gt; PRT

&lt;213&gt; Conus ebraceus

&lt;220&gt;

&lt;221&gt; PEPTIDE

&lt;222&gt; (1)..(15)

<223> Xaa at residue 3 is Glu or gamma-carboxy Glu; Xaa at residue 5 and 12 is Pro or Hyp; Xaa at residue 7 is Tyr, 125I-Tyr, mono-iodo-Tyr, di-iodo-Tyr, O-sulpho-Tyr or O-phospho-Ty

&lt;400&gt; 304

Cys	Cys	Xaa	Gln	Xaa	Cys	Xaa	Met	Gly	Cys	Ile	Xaa	Cys	Cys	Phe
1				5				10				15		

&lt;210&gt; 305

&lt;211&gt; 241

&lt;212&gt; DNA

&lt;213&gt; Conus ebraceus

&lt;400&gt; 305

ggatccatga tgtctaaact gggagtcttg ttgaccatct gtctgcttct gtttcccctt 60

actgctcttc cactggatga agatcaacct gcagacctac ctgcagagcg tatgcaggac 120

attgcaactg aacagcatcc cttgtttgat cctgtcaaac ggtgctgcgc gcagccatgc 180

tacatggat gcatcccttg ttgcttctaa taataacgtg ttgatgacca actttctcga 240

g 241

&lt;210&gt; 306

&lt;211&gt; 67

&lt;212&gt; PRT

&lt;213&gt; Conus ebraceus

&lt;400&gt; 306

Met	Met	Ser	Lys	Leu	Gly	Val	Leu	Leu	Thr	Ile	Cys	Leu	Leu	Phe
1				5				10				15		

Pro	Leu	Thr	Ala	Leu	Pro	Leu	Asp	Glu	Asp	Gln	Pro	Ala	Asp	Leu	Pro
	20				25						30				

Ala	Glu	Arg	Met	Gln	Asp	Ile	Ala	Thr	Glu	Gln	His	Pro	Leu	Phe	Asp
	35				40						45				

Pro	Val	Lys	Arg	Cys	Cys	Ala	Gln	Pro	Cys	Tyr	Met	Gly	Cys	Ile	Pro
-----	-----	-----	-----	-----	-----	-----	-----	-----	-----	-----	-----	-----	-----	-----	-----

50

55

60

Cys Cys Phe  
65

<210> 307  
<211> 15  
<212> PRT  
<213> Conus ebraceus  
  
<220>  
<221> PEPTIDE  
<222> (1)..(15)  
<223> Xaa at residue 5 and 12 is Pro or Hyp; Xaa at residue 7 is Tyr, 1  
25I-Tyr, mono-iodo-Tyr, di-iodo-Tyr, O-sulpho-Tyr or O-phospho-Ty

<400> 307  
Cys Cys Ala Gln Xaa Cys Xaa Met Gly Cys Ile Xaa Cys Cys Phe  
1 5 10 15

<210> 308  
<211> 238  
<212> DNA  
<213> Conus flavidus

<400> 308  
ggatccatga tgtctaaact gggagtccttgg ttgaccatct gtctgcttct gtttccccctt 60  
actgctgttc cgttggatgg agatcaacct gcagaccagc ctgcagagcg tatgcagaac 120  
gagcagcatc ccttggatgg tcagaaaaga aggtgctgcc ggtggccatg ccccagtata 180  
tgcggcatgg ctaggtgttg ctgcgtcatga taacgtgttg atgaccaact ttctcgag 238

<210> 309  
<211> 67  
<212> PRT  
<213> Conus flavidus

<400> 309  
Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
1 5 10 15

Pro Leu Thr Ala Val Pro Leu Asp Gly Asp Gln Pro Ala Asp Gln Pro  
20 25 30

Ala Glu Arg Met Gln Asn Glu Gln His Pro Leu Phe Asp Gln Lys Arg  
35 40 45

Arg Cys Cys Arg Trp Pro Cys Pro Ser Ile Cys Gly Met Ala Arg Cys  
50 55 60

Cys Ser Ser  
65

<210> 310  
<211> 19  
<212> PRT  
<213> Conus flavidus  
  
<220>  
<221> PEPTIDE  
<222> (1)..(19)  
<223> Xaa at residue 6 and 8 is Pro or Hyp; Xaa at residue 5 is Trp or

bromo-Tr

<400> 310  
 Arg Cys Cys Arg Xaa Xaa Cys Xaa Ser Ile Cys Gly Met Ala Arg Cys  
 1 5 10 15

Cys Ser Ser

<210> 311  
 <211> 245  
 <212> DNA  
 <213> Conus miliaris

<220>  
 <221> misc\_feature  
 <222> (1)..(245)  
 <223> n may be any nucleotide

<400> 311  
 ggatccatga tgtctaaact gggagtcttg ttgaccatct gtctgcttct gtttccaatt 60  
 actgcccttc cactggatga agatcaacct gcagaccgac ctgcagagcg tatgcaggac 120  
 attgcaactg aacagcatcc cttgtttgat cccgtcaaac ggtgttgcga ttggccatgc 180  
 agcgcaggat gctacccttg ttgcttccct taataacgtg ttgatgacca actnangnaa 240  
 aaaaaa 245

<210> 312  
 <211> 68  
 <212> PRT  
 <213> Conus miliaris

<400> 312  
 Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15

Pro Ile Thr Ala Leu Pro Leu Asp Glu Asp Gln Pro Ala Asp Arg Pro  
 20 25 30

Ala Glu Arg Met Gln Asp Ile Ala Thr Glu Gln His Pro Leu Phe Asp  
 35 40 45

Pro Val Lys Arg Cys Cys Asp Trp Pro Cys Ser Ala Gly Cys Tyr Pro  
 50 55 60

Cys Cys Phe Pro  
 65

<210> 313  
 <211> 16  
 <212> PRT  
 <213> Conus miliaris

<220>  
 <221> PEPTIDE  
 <222> (1)..(16)  
 <223> Xaa at residue 5, 12 and 16 is Pro or Hyp; Xaa at residue 4 is Trp or bromo-Trp; Xaa at residue 11 is Tyr, 125I-Tyr, mono-iodo-Tyr, di-iodo-Tyr, O-sulpho-Tyr or O-phospho-Ty

<400> 313  
 Cys Cys Asp Xaa Xaa Cys Ser Ala Gly Cys Xaa Xaa Cys Cys Phe Xaa  
 1 5 10 15

<210> 314  
 <211> 230  
 <212> DNA  
 <213> *Conus miliaris*

<220>  
 <221> misc\_feature  
 <222> (1)..(230)  
 <223> n may be any nucleotide  
 <400> 314  
 ggatccatga tgtctaaact gggagtggtg ccattcgctt ttctggctt gtttcccctg 60  
 gcaacactcc aactggatgc agatcaaccc gcagaccgac ctgcgcgtaa aaagggcatt 120  
 gcaactaaac ggcatccctt gtctgatcct gtcagagggtt gttgccctcc aatgtgcaca 180  
 ccatgcttcc cttgctgttt tcgttaataa cgtgttgatg natgatgnan 230

<210> 315  
 <211> 66  
 <212> PRT  
 <213> *Conus miliaris*

<400> 315  
 Met Met Ser Lys Leu Gly Val Val Pro Phe Val Phe Leu Val Leu Phe  
 1 5 10 15

Pro Leu Ala Thr Leu Gln Leu Asp Ala Asp Gln Pro Ala Asp Arg Pro  
 20 25 30

Ala Arg Lys Lys Gly Ile Ala Thr Lys Arg His Pro Leu Ser Asp Pro  
 35 40 45

Val Arg Gly Cys Cys Pro Pro Met Cys Thr Pro Cys Phe Pro Cys Cys  
 50 55 60

Phe Arg  
 65

<210> 316  
 <211> 16  
 <212> PRT  
 <213> *Conus miliaris*

<220>  
 <221> PEPTIDE  
 <222> (1)..(16)  
 <223> Xaa at residue 4, 9 and 12 is Pro or Hyp; Xaa at residue 5 is Tyr  
     , 125I-Tyr, mono-iodo-Tyr, di-iodo-Tyr, O-sulpho-Tyr or O-phospho  
     -Ty

<400> 316  
 Gly Cys Cys Xaa Xaa Met Cys Thr Xaa Cys Phe Xaa Cys Cys Cys Phe Arg  
 1 5 10 15

<210> 317  
 <211> 295  
 <212> DNA  
 <213> *Conus ammiralis*

<400> 317  
 caagaggat cgatacgagt tcatgatgtc taaaactggga gtcttggat ccatctgtct 60  
 gcttctgttt ccccttactg ctcttccgtt ggatggatg caacactgcag accaagctgc 120

agagcgtatg caggccgagc agcatccctt gtttgcgtaa aacacgggt gttgcaggtt 180  
 tccatccccca gatacttgca gacatttgta ttgcgggtga tgataacgtg ctgatgaccc 240  
 actttgtcat cacggctacg tcaagtgtct aatgaataag taaaatgatt gcagt 295  
 <210> 318  
 <211> 65  
 <212> PRT  
 <213> Conus ammiralis  
  
 <400> 318  
 Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15  
  
 Pro Leu Thr Ala Leu Pro Leu Asp Gly Asp Gln Pro Ala Asp Gln Ala  
 20 25 30  
  
 Ala Glu Arg Met Gln Ala Glu Gln His Pro Leu Phe Asp Gln Lys Arg  
 35 40 45  
  
 Arg Cys Cys Arg Phe Pro Cys Pro Asp Thr Cys Arg His Leu Cys Cys  
 50 55 60  
  
 Gly  
 65  
  
 <210> 319  
 <211> 16  
 <212> PRT  
 <213> Conus ammiralis  
  
 <220>  
 <221> PEPTIDE  
 <222> (1)..(16)  
 <223> Xaa at residue 6 and 8 is Pro or Hyp  
  
 <400> 319  
 Arg Cys Cys Arg Phe Xaa Cys Xaa Asp Thr Cys Arg His Leu Cys Cys  
 1 5 10 15  
  
 <210> 320  
 <211> 267  
 <212> DNA  
 <213> Conus ammiralis  
  
 <400> 320  
 caagagggtt cgatagcgt tcgtatgttt taaaactgggtt gtcttgcttgc ccatctgtct 60  
 acttctgtttt tcccttaatgtt ctgttccgtt ggatggagat caacactgc accaaccgtc 120  
 agagcgtctg ctggacgaca tttcatctgtt aaataatccc ttttatgttc cccggccaaacgt 180  
 gtgttgcattt acttgcttcgtt gttgcacacc ttgttgcattt tgaccaggctt catcaagtgt 240  
 ctaacgtttt agtaaaacgtt ttgcagt 267  
  
 <210> 321  
 <211> 66  
 <212> PRT  
 <213> Conus ammiralis  
  
 <400> 321  
 Met Met Phe Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15

Ser Leu Asn Ala Val Pro Leu Asp Gly Asp Gln Pro Ala Asp Gln Pro  
 20 25 30

Ala Glu Arg Leu Leu Asp Asp Ile Ser Ser Glu Asn Asn Pro Phe Tyr  
 35 40 45

Asp Pro Ala Lys Arg Cys Cys Met Thr Cys Phe Gly Cys Thr Pro Cys  
 50 55 60

Cys Gly  
 65

<210> 322  
 <211> 12  
 <212> PRT  
 <213> Conus ammiralis

<220>  
 <221> PEPTIDE  
 <222> (1)..(12)  
 <223> Xaa at residue 10 is Pro or Hyp

<400> 322  
 Cys Cys Met Thr Cys Phe Gly Cys Thr Xaa Cys Cys  
 1 5 10

<210> 323  
 <211> 294  
 <212> DNA  
 <213> Conus ammiralis

<400> 323  
 caagaaggat cgatacgagt tcatgatgtc taaaactggga gccttggta ccatctgtct 60  
 acttctgtt tcccttactg ctgttccgtt ggtatggagat caacatgcag accaacctgc 120  
 agagcgtctg caggaccgccc ttccaaactga aaatcatccc ttatatgatc ccgtcaaacg 180  
 gtgttgcgtt gattcggaat gcgactattc ttgctggct tgctgtatcc tttcataacc 240  
 tttgttatcg cggcctcatc ctatgtcaa atgaataagt aaaacgattt cagt 294

<210> 324  
 <211> 71  
 <212> PRT  
 <213> Conus ammiralis

<400> 324  
 Met Met Ser Lys Leu Gly Ala Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15

Ser Leu Thr Ala Val Pro Leu Asp Gly Asp Gln His Ala Asp Gln Pro  
 20 25 30

Ala Glu Arg Leu Gln Asp Arg Leu Pro Thr Glu Asn His Pro Leu Tyr  
 35 40 45

Asp Pro Val Lys Arg Cys Cys Asp Asp Ser Glu Cys Asp Tyr Ser Cys  
 50 55 60

Trp Pro Cys Cys Ile Phe Ser  
 65 70  
 <210> 325  
 <211> 18

<212> PRT  
 <213> Conus ammiralis

<220>  
 <221> PEPTIDE  
 <222> (1)..(18)

<223> Xaa at residue 6 is Glu or gamma-carboxy Glu; Xaa at residue 13 is Pro or Hyp; Xaa at residue 12 is Trp or bromo-Trp; Xaa at residue 9 is Tyr, 125I-Tyr, mono-iodo-Tyr, di-iodo-Tyr, O-sulpho-Tyr or O-phospho-Tyr

<400> 325

Cys Cys Asp Asp Ser Xaa Cys Asp Xaa Ser Cys Xaa Xaa Cys Cys Ile  
 1 5 10 15

Phe Ser

<210> 326

<211> 284

<212> DNA

<213> Conus ammiralis

<400> 326

caagagggtat cgatagcagt tcatgatgtt taaaactcggga gtcttgctga ccatctgtct 60  
 acttctgttt tccctaattt ctgttccgct ggatggagat caacatgcag accaacacctgc 120  
 agagcgtctg caggaccgccc ttccaaactga aaatcatcccc ttatatgatc ccgtcaaaacg 180  
 gtgttgcagg ttgttatgcc tcagttgcaa cccttgggttggatgaccag ctttggttatc 240  
 acggccatca caagtgtctta atgaaataagt aaaacgattt cagt 284

<210> 327

<211> 67

<212> PRT

<213> Conus ammiralis

<400> 327

Met Met Phe Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15

Ser Leu Ile Ala Val Pro Leu Asp Gly Asp Gln His Ala Asp Gln Pro  
 20 25 30

Ala Glu Arg Leu Gln Asp Arg Leu Pro Thr Glu Asn His Pro Leu Tyr  
 35 40 45

Asp Pro Val Lys Arg Cys Cys Arg Leu Leu Cys Leu Ser Cys Asn Pro  
 50 55 60

Cys Cys Gly  
 65

<210> 328

<211> 13

<212> PRT

<213> Conus ammiralis

<220>

<221> PEPTIDE

<222> (1)..(13)

<223> Xaa at residue 11 is Pro or Hyp

<400> 328

Cys Cys Arg Leu Leu Cys Leu Ser Cys Asn Xaa Cys Cys  
 1 5 10

<210> 329

<211> 289

<212> DNA

<213> Conus ammiralis

<400> 329

caagaaggat cgatagcagt tcatgatgtc taaaactggga gccttggta ccatctgtct 60

acttctgttt tcccttactg ctgttccgct ggatggagat caacatgcag accaacacctc 120

agagcgtctg caggaccgca ttccaaactga agatcatccc ttatttgcata ccaacaaacg 180

gtgttgcgtat gattcggat gcggcttattc atgctggct tgctgttatg gataagcttt 240

gttatcgcgg cctcatccag tgtcaacgaa taagtaaaac gattgcagt 289

<210> 330

<211> 70

<212> PRT

<213> Conus ammiralis

<400> 330

Met Met Ser Lys Leu Gly Ala Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15

Ser Leu Thr Ala Val Pro Leu Asp Gly Asp Gln His Ala Asp Gln Pro  
 20 25 30

Ala Glu Arg Leu Gln Asp Arg Ile Pro Thr Glu Asp His Pro Leu Phe  
 35 40 45

Asp Pro Asn Lys Arg Cys Cys Asp Asp Ser Glu Cys Gly Tyr Ser Cys  
 50 55 60

Trp Pro Cys Cys Tyr Gly  
 65 70

<210> 331

<211> 16

<212> PRT

<213> Conus ammiralis

<220>

<221> PEPTIDE

<222> (1)..(16)

<223> Xaa at residue 6 is Glu or gamma-carboxy Glu; Xaa at residue 13 is Pro or Hyp; Xaa at residue 12 is Trp or bromo-Trp; Xaa at residue 9 and 16 is Tyr, 125I-Tyr, mono-iodo-Tyr, di-iodo-Tyr, O-sulpho-Tyr or O-phospho-Ty

<400> 331

Cys Cys Asp Asp Ser Xaa Cys Gly Xaa Ser Cys Xaa Xaa Cys Cys Xaa  
 1 5 10 15

<210> 332

<211> 272

<212> DNA

<213> Conus spurius

<400> 332

caagaaggat cgatagcagt tcatgatgtc taaaactggga gtcttgcgtat ccatctgtct 60

gcttctgttt ccacgtactt ctcttccgct ggatggagat caacctgcag tccgatctgc 120  
 aaagcgatcg cattcatcta tacagcgatcg tttctttgat cccgtcaaac ggtgttgc 180  
 tagatgcagc gagtgcacccttggatgaccagcttgcgc ggcctcatta 240  
 agtgtctaat gaataagtaa aatgattgca gt 272  
  
 <210> 333  
 <211> 63  
 <212> PRT  
 <213> Conus spurius  
  
 <400> 333  
 Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15  
  
 Pro Arg Thr Ser Leu Pro Leu Asp Gly Asp Gln Pro Ala Val Arg Ser  
 20 25 30  
  
 Ala Lys Arg Met His Ser Ser Ile Gln Arg Arg Phe Phe Asp Pro Val  
 35 40 45  
  
 Lys Arg Cys Cys Pro Arg Cys Ser Glu Cys Asn Pro Cys Cys Gly  
 50 55 60  
  
 <210> 334  
 <211> 12  
 <212> PRT  
 <213> Conus spurius  
  
 <220>  
 <221> PEPTIDE  
 <222> (1)..(12)  
 <223> Xaa at residue 7 is Glu or gamma-carboxy Glu; Xaa at residue 3 and 10 is Pro or Hy  
  
 <400> 334  
 Cys Cys Xaa Arg Cys Ser Xaa Cys Asn Xaa Cys Cys  
 1 5 10  
  
 <210> 335  
 <211> 293  
 <212> DNA  
 <213> Conus omaria  
  
 <400> 335  
 caagaggat cgatagcagt tcatgatgtc taaaactggga gtctcggtga ccatctgtct 60  
 acttctatccatccgtctg ctgttccgct tgatggagat caacatgcag accaaccgtgc 120  
 agagcgatcg cagggcgaca ttttatctga aaagcatccc ttatattaatc ccgtcaaacg 180  
 gtgttgcgt gaggagaat gcagcgtgc atgctggct tggatggg ggtgtatc 240  
 tttgttatcg cggcctcatc aagtgtctaa tgaataagta aaatgattgc agt 293  
  
 <210> 336  
 <211> 70  
 <212> PRT  
 <213> Conus omaria  
  
 <400> 336  
 Met Met Ser Lys Leu Gly Val Ser Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15

Ser Leu Thr Ala Val Pro Leu Asp Gly Asp Gln His Ala Asp Gln Pro  
 20 25 30

Ala Glu Arg Leu Gln Gly Asp Ile Leu Ser Glu Lys His Pro Leu Phe  
 35 40 45

Asn Pro Val Lys Arg Cys Cys Asp Glu Glu Glu Cys Ser Ser Ala Cys  
 50 55 60

Trp Pro Cys Cys Trp Gly  
 65 70

<210> 337

<211> 16

<212> PRT

<213> Conus omaria

<220>

<221> PEPTIDE

<222> (1)..(16)

<223> Xaa at residue 4, 5 and 6 is Glu or gamma-carboxy Glu; Xaa at res  
 idue 13 is Pro or Hyp; Xaa at residue 12 and 16 is Trp or bromo-T  
 r

<400> 337

Cys Cys Asp Xaa Xaa Xaa Cys Ser Ser Ala Cys Xaa Xaa Cys Cys Xaa  
 1 5 10 15

<210> 338

<211> 293

<212> DNA

<213> Conus omaria

<400> 338

caagaaggat cgatacgagt tcatgatgtc taaaactggga gtcttgttga tcatctgtct 60

acttctgtgt ccccttactg ctgttctgga ggatggagat caacctgcag accgacacctgc 120

agagcgtatg caggacgaca tttcaactga gcatcatccc ttttatgatc ccgtcaaacg 180

gtgttgcaag tacgggtgga catgcttgct aggatgcact ccttgcattt gttgaccagt 240

tttgcattatcg cggcctcgtc aagtgtctaa tgaataagta aaacgattgc agt 293

<210> 339

<211> 70

<212> PRT

<213> Conus omaria

<400> 339

Met Met Ser Lys Leu Gly Val Leu Leu Ile Ile Cys Leu Leu Cys  
 1 5 10 15

Pro Leu Thr Ala Val Leu Glu Asp Gly Asp Gln Pro Ala Asp Arg Pro  
 20 25 30

Ala Glu Arg Met Gln Asp Asp Ile Ser Thr Glu His His Pro Phe Tyr  
 35 40 45

Asp Pro Val Lys Arg Cys Cys Lys Tyr Gly Trp Thr Cys Leu Leu Gly  
 50 55 60

Cys Thr Pro Cys Asp Cys  
 65 70

<210> 340  
 <211> 17  
 <212> PRT  
 <213> Conus omaria

<220>  
 <221> PEPTIDE  
 <222> (1)..(17)  
 <223> Xaa at residue 14 Pro or Hyp; Xaa at residue 6 is Trp or bromo-Trp; Xaa at residue 4 is Tyr, 125I-Tyr, mono-iodo-Tyr, di-iodo-Tyr, O-sulpho-Tyr or O-phospho-Tyr

<400> 340  
 Cys Cys Lys Xaa Gly Xaa Thr Cys Leu Leu Gly Cys Thr Xaa Cys Asp  
 1 5 10 15

Cys

<210> 341  
 <211> 290  
 <212> DNA  
 <213> Conus omaria

<400> 341  
 caagaggat cgatagcagt tcatgatgtc tatactggga gtcttgttga tcatctgtct 60  
 acttctgtgt ccccttactg ctgttctgga ggatggagat caacctgcag accgacctgc 120  
 agagcgtatg caggacggca tttcatctga acatcatccc tttttggatc ccgtcaaacg 180  
 gtgttgcatt ctattggcat gccgcatttgg atgctgcct tgggttgggt gaccagcttt 240  
 gttatcgccg cctcatcaag tgtctaatga ataagtaaaa cgattgcagt 290

<210> 342  
 <211> 69  
 <212> PRT  
 <213> Conus omaria

<400> 342  
 Met Met Ser Ile Leu Gly Val Leu Leu Ile Ile Cys Leu Leu Leu Cys  
 1 5 10 15  
 Pro Leu Thr Ala Val Leu Glu Asp Gly Asp Gln Pro Ala Asp Arg Pro  
 20 25 30

Ala Glu Arg Met Gln Asp Gly Ile Ser Ser Glu His His Pro Phe Leu  
 35 40 45

Asp Pro Val Lys Arg Cys Cys His Leu Leu Ala Cys Arg Phe Gly Cys  
 50 55 60

Ser Pro Cys Cys Trp  
 65

<210> 343  
 <211> 16  
 <212> PRT  
 <213> Conus omaria

<220>  
 <221> PEPTIDE  
 <222> (1)..(16)  
 <223> Xaa at residue 13 is Pro or Hyp; Xaa at residue 16 is Trp or bromo-Trp

<400> 343  
 Cys Cys His Leu Leu Ala Cys Arg Phe Gly Cys Ser Xaa Cys Cys Xaa  
 1 5 10 15

<210> 344  
 <211> 293  
 <212> DNA  
 <213> Conus omaria

<400> 344  
 caagaaggat cgatagcagt tcatgatgtc taaaactggga gtcttggta tcatctgtct 60  
 acttcttgc ccccttactg ctgttccgca ggatggagat caacctgcag accgacacctgc 120  
 agagcgtatg caggccggca tttcatctga acatcatccc ttttttgcgc ccgtcaaacg 180  
 gtgttgcagg tacgggtgga catgctggct aggatgcact ccctgtggtt gttgaccagc 240  
 tttgttatcg cggcctcatc aagtgtctaa tgaataagta aaacgattgc agt 293

<210> 345  
 <211> 70  
 <212> PRT  
 <213> Conus omaria

<400> 345  
 Met Met Ser Lys Leu Gly Val Leu Leu Ile Ile Cys Leu Leu Leu Cys  
 1 5 10 15

Pro Leu Thr Ala Val Pro Gln Asp Gly Asp Gln Pro Ala Asp Arg Pro  
 20 25 30

Ala Glu Arg Met Gln Gly Gly Ile Ser Ser Glu His His Pro Phe Phe  
 35 40 45

Asp Pro Val Lys Arg Cys Cys Arg Tyr Gly Trp Thr Cys Trp Leu Gly  
 50 55 60

Cys Thr Pro Cys Gly Cys  
 65 70

<210> 346  
 <211> 17  
 <212> PRT  
 <213> Conus omaria

<220>  
 <221> PEPTIDE  
 <222> (1)...(17)  
 <223> Xaa at residue 14 is Pro or Hyp; Xaa at residue 6 and 9 is Trp or  
 bromo-Trp; Xaa at residue 4 is Tyr, 125I-Tyr, mono-iodo-Tyr, di-  
 iodo-Tyr, O-sulpho-Tyr or O-phospho-Ty

<400> 346  
 Cys Cys Arg Xaa Gly Xaa Thr Cys Xaa Leu Gly Cys Thr Xaa Cys Gly  
 1 5 10 15

Cys

<210> 347  
 <211> 293  
 <212> DNA  
 <213> Conus episcopatus

<400> 347  
 caagaaggat cgatagcagt tcatgatgtc taaactggga gtcttgtga ccatctgtct 60  
 acttctgttt tcccttattt ctgttccgct tcatggagat caacatgcag accaacctgc 120  
 agagcgtctg cagggcgaca ttttatctga aaagcatccc ttatttatgc ctgtcaaacg 180  
 gtgttgcgt gaggacgaat gcaacagttc atgctggct ttttgggg ggtgatcagc 240  
 tttgttatcg cggcctgatc aagtgtataa tgaataagta aaacgattgc agt 293  
  
 <210> 348  
 <211> 70  
 <212> PRT  
 <213> *Conus episcopatus*  
  
 <400> 348  
 Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15  
  
 Ser Leu Ile Ala Val Pro Leu Asp Gly Asp Gln His Ala Asp Gln Pro  
 20 25 30  
  
 Ala Glu Arg Leu Gln Gly Asp Ile Leu Ser Glu Lys His Pro Leu Phe  
 35 40 45  
  
 Met Pro Val Lys Arg Cys Cys Asp Glu Asp Glu Cys Asn Ser Ser Cys  
 50 55 60  
  
 Trp Pro Cys Cys Trp Gly  
 65 70  
  
 <210> 349  
 <211> 16  
 <212> PRT  
 <213> *Conus episcopatus*  
  
 <220>  
 <221> PEPTIDE  
 <222> (1)..(16)  
 <223> Xaa at residue 4 and 6 is Glu or gamma-carboxy Glu; Xaa at residue 13 is Pro or Hyp; Xaa at residue 12 and 16 is Trp or bromo-Trp  
  
 <400> 349  
 Cys Cys Asp Xaa Asp Xaa Cys Asn Ser Ser Cys Xaa Xaa Cys Cys Xaa  
 1 5 10 15  
  
 <210> 350  
 <211> 293  
 <212> DNA  
 <213> *Conus episcopatus*  
  
 <400> 350  
 caagagggat cgatagcagt tcatgatgtc taaactggga gtcttgtga ccatctgtct 60  
 acttctgttt tcccttattt ctgttccgct tcatggagat caacatgcag accaacctgc 120  
 agagcgtctg cagggcgaca ttttatctga aaagcatccc ttatttatgc ctgtcaaacg 180  
 gtgttgcgt gaggacgaat gcagcagttc atgctggct ttttgggg gatgagcagc 240  
 tttgttatcg cggcctcatac aagtgtctaa tgaataagta aaacgattgc agt 293  
  
 <210> 351  
 <211> 70

100

&lt;212&gt; PRT

&lt;213&gt; Conus episcopatus

&lt;400&gt; 351

Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Phe  
1 5 10 15Ser Leu Ile Ala Val Pro Leu Asp Gly Asp Gln His Ala Asp Gln Pro  
20 25 30Ala Glu Arg Leu Gln Gly Asp Ile Leu Ser Glu Lys His Pro Leu Phe  
35 40 45Met Pro Val Lys Arg Cys Cys Asp Glu Asp Glu Cys Ser Ser Ser Cys  
50 55 60Trp Pro Cys Cys Trp Gly  
65 70

&lt;210&gt; 352

&lt;211&gt; 16

&lt;212&gt; PRT

&lt;213&gt; Conus episcopatus

&lt;220&gt;

&lt;221&gt; PEPTIDE

&lt;222&gt; (1)..(16)

&lt;223&gt; Xaa at residue 4 and 6 is Glu or gamma-carboxy Glu; Xaa at residue 13 is Pro or Hyp; Xaa at residue 12 and 16 is Trp or bromo-Trp

&lt;400&gt; 352

Cys Cys Asp Xaa Asp Xaa Cys Ser Ser Ser Cys Xaa Xaa Cys Cys Xaa  
1 5 10 15

&lt;210&gt; 353

&lt;211&gt; 290

&lt;212&gt; DNA

&lt;213&gt; Conus episcopatus

&lt;400&gt; 353

caagaggat cgatacgagt tcatgatgtc taaaactggga gtcttggta ccatctgtct 60

acttctgttt tcccttactg ctgttccgct tgatggagat caacatgcag accaacacctgc 120

agagcgtctg cagggcgaca ttttatctga aaagcatccc ttatthaatc ccgtcaaacg 180

gtgttgcggcg gccccatgtccatggg atgcaaggct tggatggat gagcagcttt 240

gttatcgtgg cctcatcaag tgtctaatga ataagtaaaa cgattgcagt 290

&lt;210&gt; 354

&lt;211&gt; 69

&lt;212&gt; PRT

&lt;213&gt; Conus episcopatus

&lt;400&gt; 354

Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Phe  
1 5 10 15Ser Leu Thr Ala Val Pro Leu Asp Gly Asp Gln His Ala Asp Gln Pro  
20 25 30Ala Glu Arg Leu Gln Gly Asp Ile Leu Ser Glu Lys His Pro Leu Phe  
35 40 45

101

Asn	Pro	Val	Lys	Arg	Cys	Cys	Pro	Ala	Ala	Ala	Cys	Ala	Met	Gly	Cys
50					55						60				

Lys	Pro	Cys	Cys	Gly
65				

<210>	355
<211>	15
<212>	PRT
<213>	Conus episcopatus

<220>	
<221>	PEPTIDE
<222>	(1)...(15)
<223>	Xaa at residue 3 and 13 is Pro or Hyp

<400>	355													
Cys	Cys	Xaa	Ala	Ala	Ala	Cys	Ala	Met	Gly	Cys	Lys	Xaa	Cys	Cys
1				5				10				15		

<210>	356
<211>	295
<212>	DNA
<213>	Conus aulicus

<400>	356					
caagagggtt	cgatagcagt	tcatgtatgtc	taaaactggga	gtcttggta	ccatctgtct	60
gcttctgttt		tccgttactg	ctcttccgccc	ggatggagat	caacctgcag	120
agagcgtagg	caggctcgagc	agcatcccgt	gtttgatcat	gaaagagggt	gttgctcgcc	180
accatgccac	agtatttgcg	ctgctttctg	ttgcgggtga	tgataacgtg	ttgatgacc	240
actttgtcat	cacggctcg	tcaagtgtct	aatgaataag	taaaatgatt	gcagt	295

<210>	357
<211>	65
<212>	PRT
<213>	Conus aulicus

<400>	357													
Met	Met	Ser	Lys	Leu	Gly	Val	Leu	Leu	Thr	Ile	Cys	Leu	Leu	Phe
1						5		10				15		

Ser	Val	Thr	Ala	Leu	Pro	Pro	Asp	Gly	Asp	Gln	Pro	Ala	Asp	Arg	Ala
				20				25			30				

Ala	Glu	Arg	Arg	Gln	Val	Glu	Gln	His	Pro	Val	Phe	Asp	His	Glu	Arg
				35		40					45				

Gly	Cys	Cys	Ser	Pro	Pro	Cys	His	Ser	Ile	Cys	Ala	Ala	Phe	Cys	Cys
				50		55				60					

Gly		
65		
<210>	358	
<211>	16	
<212>	PRT	
<213>	Conus aulicus	

<220>	
<221>	PEPTIDE
<222>	(1)...(16)
<223>	Xaa at residue 5 and 6 is Pro or Hyp

<400> 358  
 Gly Cys Cys Ser Xaa Xaa Cys His Ser Ile Cys Ala Ala Phe Cys Cys  
 1 5 10 15

<210> 359  
 <211> 290  
 <212> DNA  
 <213> Conus aulicus

<400> 359  
 caagagggat cgatagcagt tcatgatgtc taaaactggga gtcttggat ccatctgtct 60  
 acttctgttt tcccttactg ctgttccgct tggatggagat caacatgcag accaaccctgc 120  
 agagcgtctg cagggcgaca ttttatctga aaagcatccc ttatattaatc ccgtcaaacg 180  
 gtgttggcga ccgggtggcat gtgccatggg atgcaagcct tggatggat gaggcagctt 240  
 gttatcgtgg cctcatcaag tgtctaatga ataagtaaaa tgattgcagt 290

<210> 360  
 <211> 69  
 <212> PRT  
 <213> Conus aulicus

<400> 360  
 Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15

Ser Leu Thr Ala Val Pro Leu Asp Gly Asp Gln His Ala Asp Gln Pro  
 20 25 30

Ala Glu Arg Leu Gln Gly Asp Ile Leu Ser Glu Lys His Pro Leu Phe  
 35 40 45

Asn Pro Val Lys Arg Cys Cys Arg Pro Val Ala Cys Ala Met Gly Cys  
 50 55 60

Lys Pro Cys Cys Gly  
 65

<210> 361  
 <211> 15  
 <212> PRT  
 <213> Conus aulicus

<220>  
 <221> PEPTIDE  
 <222> (1)..(15)  
 <223> Xaa at residue 4 and 13 is Pro or Hyp

<400> 361  
 Cys Cys Arg Xaa Val Ala Cys Ala Met Gly Cys Lys Xaa Cys Cys  
 1 5 10 15

<210> 362  
 <211> 290  
 <212> DNA  
 <213> Conus aulicus

<400> 362  
 caagagggat cgatagcagt tcatgatgtc taaaactggga gtcttggat tcatctgtct 60  
 acttctgtct ccccttactg ctgttccgct ggtggagat caacatgcag accgacactgc 120

agagcgtatg caggacgaca tttcatctga acatcaaccc atgtttcatg ccatcagaca 180  
 gtgttgcggc gcggtggcat gcccattggg atgcgagcct tggatggat gaccagctt 240  
 gttatcgccg cctcatcaag tgtctaatga ataagtaaaa tgattgcagt 290  
  
 <210> 363  
 <211> 69  
 <212> PRT  
 <213> Conus aulicus  
  
 <400> 363  
 Met Met Ser Lys Leu Gly Val Leu Leu Ile Ile Cys Leu Leu Leu Ser  
 1 5 10 15  
  
 Pro Leu Thr Ala Val Pro Leu Asp Gly Asp Gln Pro Ala Asp Arg Pro  
 20 25 30  
  
 Ala Glu Arg Met Gln Asp Asp Ile Ser Ser Glu His Gln Pro Met Phe  
 35 40 45  
  
 Asp Ala Ile Arg Gln Cys Cys Pro Ala Val Ala Cys Ala Met Gly Cys  
 50 55 60  
  
 Glu Pro Cys Cys Gly  
 65  
  
 <210> 364  
 <211> 16  
 <212> PRT  
 <213> Conus aulicus  
  
 <220>  
 <221> PEPTIDE  
 <222> (1)..(16)  
 <223> Xaa at residue 1 is Gln or pyro-Glu; Xaa at residue 13 is Glu or  
 gamma-carboxy Glu; Xaa at residue 4 and 14 is Pro or Hy  
  
 <400> 364  
 Xaa Cys Cys Xaa Ala Val Ala Cys Ala Met Gly Cys Xaa Xaa Cys Cys  
 1 5 10 15  
  
 <210> 365  
 <211> 293  
 <212> DNA  
 <213> Conus aureus  
  
 <400> 365  
 caagaaggat cgatagcgt tcgtatgtc taaaactggga qccttggat ccatctgtct 60  
 acttctgttt tcccttactg ctgttccgct ggatggagat caacatgcag accaacatgc 120  
 agagcgtctg catgaccgccc ttccaaactga aaatcatccc ttatatgttc ccgtcaaacc 180  
 gtgttgcgtat gattcggat gcgactattc ttgctggct tgctgtatcc ttggataacc 240  
 tttgttatcg cggcctcatc aagtgtcaaa tgaataagta aaacgattgc agt 293  
  
 <210> 366  
 <211> 71  
 <212> PRT  
 <213> Conus aureus  
  
 <400> 366

104

Met Met Ser Lys Leu Gly Ala Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15

Ser Leu Thr Ala Val Pro Leu Asp Gly Asp Gln His Ala Asp Gln His  
 20 25 30

Ala Glu Arg Leu His Asp Arg Leu Pro Thr Glu Asn His Pro Leu Tyr  
 35 40 45

Asp Pro Val Lys Arg Cys Cys Asp Asp Ser Glu Cys Asp Tyr Ser Cys  
 50 55 60

Trp Pro Cys Cys Ile Phe Gly  
 65 70

<210> 367

<211> 17

<212> PRT

<213> Conus aureus

<220>

<221> PEPTIDE

<222> (1)..(17)

<223> Xaa at residue 6 is Glu or gamma-carboxy Glu; Xaa at residue 13 is Pro or Hyp; Xaa at residue 12 is Trp or bromo-Trp; Xaa at residue 9 is Tyr, 125I-Tyr, mono-iodo-Tyr, di-iodo-Tyr, O-sulpho-Tyr or O-phospho-Ty

<400> 367

Cys Cys Asp Asp Ser Xaa Cys Asp Xaa Ser Cys Xaa Xaa Cys Cys Ile  
 1 5 10 15

Phe

<210> 368

<211> 290

<212> DNA

<213> Conus aureus

<400> 368

caagaggat cgatagcagt tcatgatgtc taaaactggga gccttggta ccatctgtct 60

acttctgtt tccctaactg ctgttccgct ggatggagat caacatgcag accaacacctgc 120

agagcgtctg caggaccgca ttccaaactga aaatcatccc ttatgtatc cgaacaaacg 180

gtgttgcaat gattggaaat gcgacgattc atgctggcct tgctgttatg gataaccttt 240

gttatcgcgg cctcatcaag tgtcaaatga ataagtaaaa cgattgcagt 290

<210> 369

<211> 70

<212> PRT

<213> Conus aureus

<400> 369

Met Met Ser Lys Leu Gly Ala Leu Leu Thr Ile Cys Leu Leu Leu Phe  
 1 5 10 15

Ser Leu Thr Ala Val Pro Leu Asp Gly Asp Gln His Ala Asp Gln Pro  
 20 25 30

Ala Glu Arg Leu Gln Asp Arg Ile Pro Thr Glu Asn His Pro Leu Phe  
 35 40 45

105

Asp	Pro	Asn	Lys	Arg	Cys	Cys	Asn	Asp	Trp	Glu	Cys	Asp	Asp	Ser	Cys
50					55					60					

Trp	Pro	Cys	Cys	Tyr	Gly
65				70	

&lt;210&gt; 370

&lt;211&gt; 16

&lt;212&gt; PRT

&lt;213&gt; Conus aureus

&lt;220&gt;

&lt;221&gt; PEPTIDE

&lt;222&gt; (1)..(16)

<223> Xaa at residue 6 is Glu or gamma-carboxy Glu; Xaa at residue 13 is Pro or Hyp; Xaa at residue 5 and 12 is Trp or bromo-Trp; Xaa at residue 16 is Tyr, 125I-Tyr, mono-iodo-Tyr, di-iodo-Tyr, O-sulpho-Tyr or O-phospho-Tyr

&lt;400&gt; 370

Cys	Cys	Asn	Asp	Xaa	Xaa	Cys	Asp	Asp	Ser	Cys	Xaa	Xaa	Cys	Cys	Xaa
1				5					10				15		

&lt;210&gt; 371

&lt;211&gt; 310

&lt;212&gt; DNA

&lt;213&gt; Conus consors

&lt;400&gt; 371

caagagggat	cgatagcagt	tcatgatgtc	taaactggga	gtcttggta	ccatctgttt	60
gcttctgttt	cccccttactg	ctcttccat	ggatggagat	caatctgtag	accgacacctgc	120
agagcgtatg	caggacgaca	tttcatctga	gctgcattccc	ttgttcaatc	agaaaaagaat	180
gtgttgcggc	gaaggtgcgc	catgccccag	ctatttcaga	aacagtcaga	tttgcattg	240
ttgttaaatg	acaacgtgtc	gatgaccaac	ttcggttatca	cgactaatga	ataagtaaaa	300
tgattgcagt						310

&lt;210&gt; 372

&lt;211&gt; 74

&lt;212&gt; PRT

&lt;213&gt; Conus consors

&lt;400&gt; 372

Met	Met	Ser	Lys	Leu	Gly	Val	Leu	Leu	Thr	Ile	Cys	Leu	Leu	Leu	Phe
1				5					10				15		

Pro	Leu	Thr	Ala	Leu	Pro	Met	Asp	Gly	Asp	Gln	Ser	Val	Asp	Arg	Pro
				20			25			30					

Ala	Glu	Arg	Met	Gln	Asp	Asp	Ile	Ser	Ser	Glu	Leu	His	Pro	Leu	Phe
				35			40			45					

Asn	Gln	Lys	Arg	Met	Cys	Cys	Gly	Glu	Gly	Ala	Pro	Cys	Pro	Ser	Tyr
				50			55			60					

Phe	Arg	Asn	Ser	Gln	Ile	Cys	His	Cys	Cys
65				70					

&lt;210&gt; 373

&lt;211&gt; 22

&lt;212&gt; PRT

&lt;213&gt; Conus consors

&lt;220&gt;

&lt;221&gt; PEPTIDE

&lt;222&gt; (1)..(22)

&lt;223&gt; Xaa at residue 5 is Glu or gamma-carboxy Glu; Xaa at residue 8 and 10 is Pro or Hyp; Xaa at residue 12 is Tyr, 125I-Tyr, mono-iodo-Tyr, di-iodo-Tyr, O-sulpho-Tyr or O-phospho-Ty

&lt;400&gt; 373

Met Cys Cys Gly Xaa Gly Ala Xaa Cys Xaa Ser Xaa Phe Arg Asn Ser  
1 5 10 15Gln Ile Cys His Cys Cys  
20

&lt;210&gt; 374

&lt;211&gt; 315

&lt;212&gt; DNA

&lt;213&gt; Conus consors

&lt;400&gt; 374

taagaggat ctagatcgat tcatgtatgtc taaaactggga gtcttgttga ccatctgtct 60  
gcttctgttt ccccttattt ctcttccaaat ggatggagat caacctgcag accgacactgc 120  
agagcgtatg caggaacgaca tttcatctca gcagcatccc ttgtttgata agagaggccg 180  
ctgttgcgat gtgcgaacg catgctccgg cagatggtgc agagatcacy cacaatgttg 240  
cgatgacga taacgtgttg atgaccaact ttgtgatcac ggctacatca agtgaataag 300  
taaaaacgatt gcagt 315

&lt;210&gt; 375

&lt;211&gt; 74

&lt;212&gt; PRT

&lt;213&gt; Conus consors

&lt;400&gt; 375

Met Met Ser Lys Leu Gly Val Leu Leu Thr Ile Cys Leu Leu Leu Phe  
1 5 10 15Pro Leu Ile Ala Leu Pro Met Asp Gly Asp Gln Pro Ala Asp Arg Pro  
20 25 30Ala Glu Arg Met Gln Asp Asp Ile Ser Ser Gln Gln His Pro Leu Phe  
35 40 45Asp Lys Arg Gly Arg Cys Cys Asp Val Pro Asn Ala Cys Ser Gly Arg  
50 55 60Trp Cys Arg Asp His Ala Gln Cys Cys Gly  
65 70

&lt;210&gt; 376

&lt;211&gt; 22

&lt;212&gt; PRT

&lt;213&gt; Conus consors

&lt;220&gt;

&lt;221&gt; PEPTIDE

&lt;222&gt; (1)..(22)

&lt;223&gt; Xaa at residue 7 is Pro or Hyp; Xaa at residue 14 is Trp or bromo-Tr

<400> 376  
 Gly Arg Cys Cys Asp Val Xaa Asn Ala Cys Ser Gly Arg Xaa Cys Arg  
 1 5 10 15

Asp His Ala Gln Cys Cys  
 20

<210> 377  
 <211> 322  
 <212> DNA  
 <213> *Conus consors*

<400> 377  
 caagaggat cgatacgagt tcatgatgtc taaaactggga gtcttgttga ctgtctgttt 60  
 gcttctgttt ccccttactg ctcttccgat ggatggagat caacctgcag accaacacctgc 120  
 agagcgtatg caggacgaca tttcatctga gcagcataccc ttgtttgata agagacaaag 180  
 gtgttgcact gggagaagg ggtcatgctc cggtaaagca tgcaaaagtc tcaaattgttg 240  
 ctctggacga taacgtgttg atgaccaact ttgttatcac ggctacgtca agtgtctagt 300  
 gaataagtaa aacgattgca gt 322

<210> 378  
 <211> 76  
 <212> PRT  
 <213> *Conus consors*

<400> 378  
 Met Met Ser Lys Leu Gly Val Leu Leu Thr Val Cys Leu Leu Leu Phe  
 1 5 10 15

Pro Leu Thr Ala Leu Pro Met Asp Gly Asp Gln Pro Ala Asp Gln Pro  
 20 25 30

Ala Glu Arg Met Gln Asp Asp Ile Ser Ser Glu Gln His Pro Leu Phe  
 35 40 45

Asp Lys Arg Gln Arg Cys Cys Thr Gly Lys Lys Gly Ser Cys Ser Gly  
 50 55 60

Lys Ala Cys Lys Ser Leu Lys Cys Cys Ser Gly Arg  
 65 70 75

<210> 379  
 <211> 23  
 <212> PRT  
 <213> *Conus consors*

<220>  
 <221> PEPTIDE  
 <222> (1)..(23)  
 <223> Xaa at residue 1 is Gln or pyro-Glu

<400> 379  
 Xaa Arg Cys Cys Thr Gly Lys Lys Gly Ser Cys Ser Gly Lys Ala Cys  
 1 5 10 15

Lys Ser Leu Lys Cys Cys Ser  
 20

<210> 380

&lt;211&gt; 284

&lt;212&gt; DNA

&lt;213&gt; Conus emaciatus

&lt;400&gt; 380

caagaggat cgatacgagt tcatgatgtc taaaactggga gtcttgctga ccatctgtct 60  
 gcttctgttt ccccttactg ttcttccgat ggatggagat caacctgcag acctacacctc 120  
 attgcgtgcg cagttctttg cacctgaaca tagtccccgg tttgaccccg tcaaacggtg 180  
 ctgctcgcgg gattgcagtg tttgcatccc ttgttgcaccc tatggatcac ctgattatt 240  
 gcgccacgt caagtgtcta atgaataagt aaaatgattt cagt 284

&lt;210&gt; 381

&lt;211&gt; 70

&lt;212&gt; PRT

&lt;213&gt; Conus emaciatus

&lt;400&gt; 381

Met	Met	Ser	Lys	Leu	Gly	Val	Leu	Leu	Thr	Ile	Cys	Leu	Leu	Leu	Phe
1				5				10					15		

Pro	Leu	Thr	Val	Leu	Pro	Met	Asp	Gly	Asp	Gln	Pro	Ala	Asp	Leu	Pro
			20			25			30						

Ala	Leu	Arg	Ala	Gln	Phe	Phe	Ala	Pro	Glu	His	Ser	Pro	Arg	Phe	Asp
					35		40			45					

Pro	Val	Lys	Arg	Cys	Cys	Ser	Arg	Asp	Cys	Ser	Val	Cys	Ile	Pro	Cys
	50				55				60						

Cys	Pro	Tyr	Gly	Ser	Pro
65			70		

&lt;210&gt; 382

&lt;211&gt; 18

&lt;212&gt; PRT

&lt;213&gt; Conus emaciatus

&lt;220&gt;

&lt;221&gt; PEPTIDE

&lt;222&gt; (1)..(18)

<223> Xaa at residue 11, 14 and 18 is Pro or Hyp; Xaa at residue 15 is Tyr, 125I-Tyr, mono-iodo-Tyr, di-iodo-Tyr, O-sulpho-Tyr or O-phospho-Ty

&lt;400&gt; 382

Cys	Cys	Ser	Arg	Asp	Cys	Ser	Val	Cys	Ile	Xaa	Cys	Cys	Xaa	Xaa	Gly
1					5			10					15		

Ser Xaa

&lt;210&gt; 383

&lt;211&gt; 13

&lt;212&gt; PRT

&lt;213&gt; Conus aurisiacus

&lt;400&gt; 383

Cys	Cys	Lys	Val	Gln	Cys	Glu	Ser	Cys	Thr	Pro	Cys	Cys
1				5				10				

&lt;210&gt; 384

&lt;211&gt; 15

&lt;212&gt; PRT

&lt;213&gt; Conus atlanticus

&lt;400&gt; 384

Cys Cys Glu Leu Pro Cys Gly Pro Gly Phe Cys Val Pro Cys Cys  
1 5 10 15

&lt;210&gt; 385

&lt;211&gt; 14

&lt;212&gt; PRT

&lt;213&gt; Conus arentus

&lt;400&gt; 385

Cys Cys Glu Arg Pro Cys Asn Ile Gly Cys Val Pro Cys Cys  
1 5 10

&lt;210&gt; 386

&lt;211&gt; 16

&lt;212&gt; PRT

&lt;213&gt; Conus bandus

&lt;400&gt; 386

Cys Cys Asn Trp Pro Cys Ser Met Gly Cys Ile Pro Cys Cys Tyr Tyr  
1 5 10 15

&lt;210&gt; 387

&lt;211&gt; 15

&lt;212&gt; PRT

&lt;213&gt; Conus betulinus

&lt;400&gt; 387

Cys Cys Glu Leu Pro Cys His Gly Cys Val Pro Cys Cys Trp Pro  
1 5 10 15

&lt;210&gt; 388

&lt;211&gt; 16

&lt;212&gt; PRT

&lt;213&gt; Conus betulinus

&lt;400&gt; 388

Cys Cys Gly Leu Pro Cys Asn Gly Cys Val Pro Cys Cys Trp Pro Ser  
1 5 10 15

&lt;210&gt; 389

&lt;211&gt; 18

&lt;212&gt; PRT

&lt;213&gt; Conus betulinus

&lt;400&gt; 389

Cys Cys Ser Arg Asn Cys Ala Val Cys Ile Pro Cys Cys Pro Asn Trp  
1 5 10 15

Pro Ala

&lt;210&gt; 390

&lt;211&gt; 14

&lt;212&gt; PRT

&lt;213&gt; Conus betulinus

&lt;400&gt; 390

Cys Cys Lys Gln Ser Cys Thr Thr Cys Met Pro Cys Cys Trp  
1 5 10

&lt;210&gt; 391

&lt;211&gt; 14

110

<212> PRT  
<213> Conus betulinus

<220>  
<221> PEPTIDE  
<222> (1)..(14)  
<223> Xaa is Glu or gamma-carboxy Glu

<400> 391  
Ala Cys Cys Xaa Gln Ser Cys Thr Thr Cys Met Pro Cys Cys  
1 5 10

<210> 392  
<211> 14  
<212> PRT  
<213> Conus betulinus

<400> 392  
Cys Cys Glu Gln Ser Cys Thr Thr Cys Met Pro Cys Cys Trp  
1 5 10

<210> 393  
<211> 18  
<212> PRT  
<213> Conus characteristicus

<400> 393  
Arg Cys Cys Arg Tyr Pro Cys Pro Asp Ser Cys His Gly Ser Cys Cys  
1 5 10 15

Tyr Lys

<210> 394  
<211> 15  
<212> PRT  
<213> Conus characteristicus

<400> 394  
Cys Cys Pro Pro Val Ala Cys Asn Met Gly Cys Lys Pro Cys Cys  
1 5 10 15

<210> 395  
<211> 17  
<212> PRT  
<213> Conus characteristicus

<400> 395  
Cys Cys Asp Asp Ser Glu Cys Asp Tyr Ser Cys Trp Pro Cys Cys Met  
1 5 10 15

Phe

<210> 396  
<211> 14  
<212> PRT  
<213> Conus characteristicus

<400> 396  
Cys Cys Arg Arg Cys Tyr Met Gly Cys Ile Pro Cys Cys Phe  
1 5 10

<210> 397  
<211> 16  
<212> PRT  
<213> Conus textile

<400> 397  
Cys Cys Pro Pro Val Ala Cys Asn Met Gly Cys Lys Pro Cys Cys Gly  
1 5 10 15

<210> 398  
<211> 19  
<212> PRT  
<213> Conus marmoreus

<220>  
<221> PEPTIDE  
<222> (1)..(19)  
<223> Xaa is Hyp

<400> 398  
Ser Lys Gln Cys Cys His Leu Ala Ala Cys Arg Phe Gly Cys Thr Xaa  
1 5 10 15

Cys Cys Asn

<210> 399  
<211> 15  
<212> PRT  
<213> Conus capitaneus

<400> 399  
Ser Cys Cys Arg Asp Cys Gly Glu Asp Cys Val Gly Cys Cys Arg  
1 5 10 15

<210> 400  
<211> 16  
<212> PRT  
<213> Conus coronatus

<400> 400  
Cys Cys Asp Trp Pro Cys Ile Pro Gly Cys Thr Pro Cys Cys Leu Pro  
1 5 10 15

<210> 401  
<211> 18  
<212> PRT  
<213> Conus dalli

<400> 401  
Cys Cys Asp Asp Ser Glu Cys Asp Tyr Ser Cys Trp Pro Cys Cys Ile  
1 5 10 15

Leu Ser

<210> 402  
<211> 17  
<212> PRT  
<213> Conus dalli

<400> 402  
Glx Gln Cys Cys Pro Pro Val Ala Cys Asn Met Gly Cys Glu Pro Cys  
1 5 10 15

Cys

<210> 403  
<211> 16  
<212> PRT  
<213> Conus dalli

<400> 403  
Cys Cys Asn Ala Gly Phe Cys Arg Phe Gly Cys Thr Pro Cys Cys Trp  
1 5 10 15

<210> 404  
<211> 14  
<212> PRT  
<213> Conus distans

<400> 404  
Glx Cys Cys Val His Pro Cys Pro Cys Thr Pro Cys Cys Arg  
1 5 10

<210> 405  
<211> 14  
<212> PRT  
<213> Conus figulinus

<400> 405  
Cys Cys Pro Trp Pro Cys Asn Ile Gly Cys Val Pro Cys Cys  
1 5 10

<210> 406  
<211> 14  
<212> PRT  
<213> Conus figulinus

<400> 406  
Cys Cys Ser Lys Asn Cys Ala Val Cys Ile Pro Cys Cys Pro  
1 5 10

<210> 407  
<211> 15  
<212> PRT  
<213> Conus figulinus

<400> 407  
Cys Cys Arg Trp Pro Cys Pro Ala Arg Cys Gly Ser Cys Cys Leu  
1 5 10 15

<210> 408  
<211> 16  
<212> PRT  
<213> Conus figulinus

<400> 408  
Cys Cys Glu Leu Ser Arg Cys Leu Gly Cys Val Pro Cys Cys Thr Ser  
1 5 10 15

<210> 409  
<211> 16  
<212> PRT  
<213> Conus figulinus

<400> 409  
Cys Cys Glu Leu Ser Lys Cys His Gly Cys Val Pro Cys Cys Ile Pro  
1 5 10 15

<210> 410  
<211> 16  
<212> PRT  
<213> Conus generalis

<400> 410

113

Glx Cys Cys Thr Phe Cys Asn Phe Gly Cys Gln Pro Cys Cys Val Pro  
1 5 10 15

<210> 411  
<211> 16  
<212> PRT  
<213> Conus generalis

<400> 411  
Glx Cys Cys Thr Phe Cys Asn Phe Gly Cys Gln Pro Cys Cys Leu Thr  
1 5 10 15

<210> 412  
<211> 16  
<212> PRT  
<213> Conus generalis

<400> 412

Glx Cys Cys Thr Phe Cys Asn Phe Gly Cys Gln Pro Cys Cys Val Pro  
1 5 10 15

<210> 413  
<211> 17  
<212> PRT  
<213> Conus gloriamaris

<400> 413  
Cys Cys Asp Asp Ser Glu Cys Asp Tyr Ser Cys Trp Pro Cys Cys Met  
1 5 10 15

Phe

<210> 414  
<211> 17  
<212> PRT  
<213> Conus gloriamaris

<400> 414  
Gly Cys Cys His Leu Leu Ala Cys Arg Phe Gly Cys Ser Pro Cys Cys  
1 5 10 15  
Trp

<210> 415  
<211> 16  
<212> PRT  
<213> Conus gloriamaris

<400> 415  
Cys Cys Ser Trp Asp Val Cys Asp His Pro Ser Cys Thr Cys Cys Gly  
1 5 10 15

<210> 416  
<211> 13  
<212> PRT  
<213> Conus laterculatus

<400> 416  
Cys Cys Asp Trp Pro Cys Ser Gly Cys Ile Pro Cys Cys  
1 5 10

<210> 417  
<211> 19  
<212> PRT  
<213> Conus leopardus

<400> 417  
Glx Ile Asn Cys Cys Pro Trp Pro Cys Pro Ser Thr Cys Arg His Gln  
1 5 10 15

Cys Cys His

<210> 418  
<211> 19  
<212> PRT  
<213> Conus lividus

<400> 418  
Glx Ile Asn Cys Cys Pro Trp Pro Cys Pro Asp Ser Cys His Tyr Gln  
1 5 10 15

Cys Cys His

<210> 419  
<211> 14  
<212> PRT  
<213> Conus marmoreus

<400> 419  
Cys Cys Arg Leu Ser Cys Gly Leu Gly Cys His Pro Cys Cys  
1 5 10

<210> 420  
<211> 17  
<212> PRT  
<213> Conus marmoreus

<400> 420  
Glu Cys Cys Gly Ser Phe Ala Cys Arg Phe Gly Cys Val Pro Cys Cys  
1 5 10 15

Val

<210> 421  
<211> 19  
<212> PRT  
<213> Conus marmoreus

<400> 421  
Ser Lys Gln Cys Cys His Leu Pro Ala Cys Arg Phe Gly Cys Thr Pro  
1 5 10 15

Cys Cys Trp

<210> 422  
<211> 17  
<212> PRT  
<213> Conus marmoreus

<400> 422  
Met Gly Cys Cys Pro Phe Pro Cys Lys Thr Ser Cys Thr Thr Leu Cys  
1 5 10 15

Cys

<210> 423  
<211> 14  
<212> PRT  
<213> Conus musicus

<400> 423

Ala Cys Cys Glu Gln Ser Cys Thr Thr Cys Phe Pro Cys Cys  
1 5 10

<210> 424  
<211> 15  
<212> PRT  
<213> Conus nobilis

<400> 424  
Cys Cys Glu Leu Pro Cys Gly Pro Gly Phe Cys Val Pro Cys Cys  
1 5 10 15

<210> 425  
<211> 14  
<212> PRT  
<213> Conus pulicarius

<400> 425  
Cys Cys Asn Ser Cys Tyr Met Gly Cys Ile Pro Cys Cys Phe  
1 5 10

<210> 426  
<211> 17  
<212> PRT  
<213> Conus quercinus

<400> 426  
Glx Arg Cys Cys Gln Trp Pro Cys Pro Gly Ser Cys Arg Cys Cys Arg  
1 5 10 15

Thr

<210> 427  
<211> 18  
<212> PRT  
<213> Conus quercinus

<400> 427  
Glx Arg Cys Cys Arg Trp Pro Cys Pro Gly Ser Cys Arg Cys Cys Arg  
1 5 10 15

Tyr Arg

<210> 428  
<211> 18  
<212> PRT  
<213> Conus quercinus

<400> 428  
Arg Cys Cys Arg Tyr Pro Cys Pro Asp Ser Cys His Gly Ser Cys Cys  
1 5 10 15

Tyr Lys

<210> 429  
<211> 15  
<212> PRT  
<213> Conus quercinus

<220>  
<221> PEPTIDE  
<222> (1)..(15)  
<223> Xaa is Hyp

<400> 429

Cys Cys Ser Gln Asp Cys Leu Val Cys Ile Xaa Cys Cys Pro Asn  
1 5 10 15

<210> 430  
<211> 15  
<212> PRT  
<213> *Conus quercinus*

<220>  
<221> PEPTIDE  
<222> (1)..(15)  
<223> Xaa is Hyp

<400> 430  
Cys Cys Ser Arg His Cys Trp Val Cys Ile Xaa Cys Cys Pro Asn  
1 5 10 15

<210> 431  
<211> 16  
<212> PRT  
<213> *Conus rattus*

<400> 431  
Glx Thr Cys Cys Ser Asn Cys Gly Glu Asp Cys Asp Gly Cys Cys Gln  
1 5 10 15

<210> 432  
<211> 20  
<212> PRT  
<213> *Conus striatus*

<400> 432  
Glx Asn Cys Cys Asn Gly Gly Cys Ser Ser Lys Trp Cys Arg Asp His  
1 5 10 15

Ala Arg Cys Cys  
20

<210> 433  
<211> 12  
<212> PRT  
<213> *Conus textile*

<220>  
<221> PEPTIDE  
<222> (1)..(12)  
<223> Xaa is Hyp

<400> 433  
Cys Cys Arg Thr Cys Phe Gly Cys Thr Xaa Cys Cys  
1 5 10

<210> 434  
<211> 14  
<212> PRT  
<213> *Conus tessulatus*

<400> 434  
Cys Cys His Lys Cys Tyr Met Gly Cys Ile Pro Cys Cys Ile  
1 5 10

<210> 435  
<211> 18  
<212> PRT  
<213> *Conus tessulatus*

<400> 435  
Lys Cys Cys Arg Pro Pro Cys Ala Met Ser Cys Gly Met Ala Arg Cys  
1 5 10 15

Cys Tyr

<210> 436  
<211> 23  
<212> PRT  
<213> Conus betulinus

<400> 436  
Arg Cys Cys Arg Trp Pro Cys Pro Ser Ile Cys Gly Met Ala Arg Cys  
1 5 10 15

Cys Phe Val Met Ile Thr Cys  
20

<210> 437  
<211> 23  
<212> PRT  
<213> Conus betulinus

<400> 437  
Arg Cys Cys Arg Trp Pro Cys Pro Ser Arg Cys Gly Met Ala Arg Cys  
1 5 10 15

Cys Phe Val Met Ile Thr Cys  
20

<210> 438  
<211> 15  
<212> PRT  
<213> Conus textile

<400> 438  
Phe Cys Cys Asp Ser Asn Trp Cys His Asp Cys Glu Cys Cys Tyr  
1 5 10 15

<210> 439  
<211> 16  
<212> PRT  
<213> Conus marmoreus

<400> 439  
Cys Cys His Trp Asn Trp Cys Asp His Leu Cys Ser Cys Cys Gly Ser  
1 5 10 15

<210> 440  
<211> 16  
<212> PRT  
<213> Conus marmoreus

<220>  
<221> PEPTIDE  
<222> (1)..(16)  
<223> Xaa is Hyp

<400> 440  
Asp Cys Cys Xaa Leu Pro Ala Cys Pro Phe Gly Cys Asn Xaa Cys Cys  
1 5 10 15

<210> 441  
<211> 16

<212> PRT  
<213> Conus marmoreus

<220>  
<221> PEPTIDE  
<222> (1)..(16)  
<223> Xaa is Hyp

<400> 441  
Cys Cys Ala Pro Ser Ala Cys Arg Leu Gly Cys Arg Xaa Cys Cys Arg  
1 5 10 15

<210> 442  
<211> 16  
<212> PRT  
<213> Conus marmoreus

<220>  
<221> PEPTIDE  
<222> (1)..(16)  
<223> Xaa is Hyp

<400> 442  
Cys Cys Ala Xaa Ser Ala Cys Arg Leu Gly Cys Arg Xaa Cys Cys Arg  
1 5 10 15

<210> 443  
<211> 16  
<212> PRT  
<213> Conus marmoreus

<400> 443  
Cys Cys Ala Pro Ser Ala Cys Arg Leu Gly Cys Arg Pro Cys Cys Arg  
1 5 10 15

<210> 444  
<211> 17  
<212> PRT  
<213> Conus marmoreus

<220>  
<221> PEPTIDE  
<222> (1)..(17)  
<223> Xaa is Hyp

<400> 444  
Gly Cys Cys Gly Ser Phe Ala Cys Arg Phe Gly Cys Val Xaa Cys Cys  
1 5 10 15

Val

<210> 445  
<211> 15  
<212> PRT  
<213> Conus textile

<400> 445  
Cys Cys Ser Trp Asp Val Cys Asp His Pro Ser Cys Thr Cys Cys  
1 5 10 15

<210> 446  
<211> 16  
<212> PRT  
<213> Conus textile

<400> 446  
Arg Cys Cys Lys Phe Pro Cys Pro Asp Ser Cys Arg Tyr Leu Cys Cys  
1 5 10 15

<210> 447  
<211> 17  
<212> PRT  
<213> Conus aureus

<400> 447  
Cys Cys Asp Asp Ser Glu Cys Asp Tyr Ser Cys Trp Pro Cys Cys Ile  
1 5 10 15

Phe

<210> 448  
<211> 16  
<212> PRT  
<213> Conus aureus

<400> 448  
Cys Cys Asn Asp Trp Glu Cys Asp Asp Ser Cys Trp Pro Cys Cys Tyr  
1 5 10 15

<210> 449  
<211> 16  
<212> PRT  
<213> Conus ammiralis

<400> 449  
Arg Cys Cys Arg Phe Pro Cys Pro Asp Thr Cys Arg His Leu Cys Cys  
1 5 10 15

<210> 450  
<211> 12  
<212> PRT  
<213> Conus ammiralis

<400> 450  
Cys Cys Met Thr Cys Phe Gly Cys Thr Pro Cys Cys  
1 5 10

<210> 451  
<211> 18  
<212> PRT  
<213> Conus ammiralis

<400> 451  
Cys Cys Asp Asp Ser Glu Cys Asp Tyr Ser Cys Trp Pro Cys Cys Ile  
1 5 10 15

Phe Ser

<210> 452  
<211> 13  
<212> PRT  
<213> Conus ammiralis

<400> 452  
Cys Cys Arg Leu Leu Cys Leu Ser Cys Asn Pro Cys Cys  
1 5 10

<210> 453  
<211> 16  
<212> PRT

120

&lt;213&gt; Conus ammiralis

&lt;400&gt; 453

Cys Cys Asp Asp Ser Glu Cys Gly Tyr Ser Cys Trp Pro Cys Cys Tyr  
1 5 10 15

&lt;210&gt; 454

&lt;211&gt; 16

&lt;212&gt; PRT

&lt;213&gt; Conus aulicus

&lt;400&gt; 454

Gly Cys Cys Ser Pro Pro Cys His Ser Ile Cys Ala Ala Phe Cys Cys  
1 5 10 15

&lt;210&gt; 455

&lt;211&gt; 15

&lt;212&gt; PRT

&lt;213&gt; Conus aulicus

&lt;400&gt; 455

Cys Cys Arg Pro Val Ala Cys Ala Met Gly Cys Lys Pro Cys Cys  
1 5 10 15

&lt;210&gt; 456

&lt;211&gt; 16

&lt;212&gt; PRT

&lt;213&gt; Conus aulicus

&lt;400&gt; 456

Glx Cys Cys Pro Ala Val Ala Cys Ala Met Gly Cys Glu Pro Cys Cys  
1 5 10 15

&lt;210&gt; 457

&lt;211&gt; 18

&lt;212&gt; PRT

&lt;213&gt; Conus emaciatus

&lt;400&gt; 457

Cys Cys Ser Arg Asp Cys Ser Val Cys Ile Pro Cys Cys Pro Tyr Gly  
1 5 10 15

Ser Pro

&lt;210&gt; 458

&lt;211&gt; 16

&lt;212&gt; PRT

&lt;213&gt; Conus episcopatus

&lt;400&gt; 458

Cys Cys Asp Glu Asp Glu Cys Asn Ser Ser Cys Trp Pro Cys Cys Trp  
1 5 10 15

&lt;210&gt; 459

&lt;211&gt; 16

&lt;212&gt; PRT

&lt;213&gt; Conus episcopatus

&lt;400&gt; 459

Cys Cys Asp Glu Asp Glu Cys Ser Ser Ser Cys Trp Pro Cys Cys Trp  
1 5 10 15

&lt;210&gt; 460

&lt;211&gt; 15

&lt;212&gt; PRT

<213> Conus episcopatus

<400> 460

Cys Cys Pro Ala Ala Ala Cys Ala Met Gly Cys Lys Pro Cys Cys  
1 5 10 15

<210> 461

<211> 16

<212> PRT

<213> Conus omaria

<400> 461

Cys Cys Asp Glu Glu Glu Cys Ser Ser Ala Cys Trp Pro Cys Cys Trp  
1 5 10 15

<210> 462

<211> 16

<212> PRT

<213> Conus omaria

<400> 462

Cys Cys His Leu Leu Ala Cys Arg Phe Gly Cys Ser Pro Cys Cys Trp  
1 5 10 15

<210> 463

<211> 12

<212> PRT

<213> Conus spurius

<400> 463

Cys Cys Pro Arg Cys Ser Glu Cys Asn Pro Cys Cys  
1 5 10

<210> 464

<211> 16

<212> PRT

<213> Conus pennaceus

<400> 464

Arg Cys Cys Lys Phe Pro Cys Pro Asp Ser Cys Lys Tyr Leu Cys Cys  
1 5 10 15

<210> 465

<211> 19

<212> PRT

<213> Conus flavidus

<400> 465

Arg Cys Cys Arg Trp Pro Cys Pro Ser Ile Cys Gly Met Ala Arg Cys  
1 5 10 15

Cys Ser Ser

<210> 466

<211> 14

<212> PRT

<213> Conus pulicarius

<400> 466

Cys Cys Lys Leu Leu Cys Gly Cys Thr Pro Cys Cys His Ile  
1 5 10

<210> 467

<211> 15

<212> PRT

<213> Conus ebraceus

<400> 467

Cys Cys Glu Gln Pro Cys Tyr Met Gly Cys Ile Pro Cys Cys Phe  
1 5 10 15

<210> 468

<211> 15

<212> PRT

<213> Conus ebraceus

<400> 468

Cys Cys Ala Gln Pro Cys Tyr Met Gly Cys Ile Pro Cys Cys Phe  
1 5 10 15

<210> 469

<211> 14

<212> PRT

<213> Conus pulicarius

<400> 469

Cys Cys Val Ser Cys Tyr Met Gly Cys Ile Pro Cys Cys Phe  
1 5 10

<210> 470

<211> 16

<212> PRT

<213> Conus miliaris

<400> 470

Cys Cys Asp Trp Pro Cys Ser Ala Gly Cys Tyr Pro Cys Cys Phe Pro  
1 5 10 15

<210> 471

<211> 16

<212> PRT

<213> Conus miliaris

<400> 471

Gly Cys Cys Pro Pro Met Cys Thr Pro Cys Phe Pro Cys Cys Phe Arg  
1 5 10 15

<210> 472

<211> 23

<212> PRT

<213> Conus rattus

<400> 472

Arg Gly Cys Cys Ala Pro Pro Arg Lys Cys Lys Asp Arg Ala Cys Lys  
1 5 10 15

Pro Ala Arg Cys Cys Gly Pro  
20

<210> 473

<211> 22

<212> PRT

<213> Conus stercusmuscarum

<400> 473

Glx Arg Cys Cys Asn Gly Arg Arg Gly Cys Ser Ser Arg Trp Cys Arg  
1 5 10 15

Asp His Ser Arg Cys Cys  
20

123

<210> 474  
<211> 22  
<212> PRT  
<213> Conus consors

<400> 474  
Gly Arg Cys Cys Asp Val Pro Asn Ala Cys Ser Gly Arg Trp Cys Arg  
1 5 10 15  
Asp His Ala Gln Cys Cys  
20

<210> 475  
<211> 23  
<212> PRT  
<213> Conus consors

<400> 475  
Glx Arg Cys Cys Thr Gly Lys Lys Gly Ser Cys Ser Gly Lys Ala Cys  
1 5 10 15  
Lys Ser Leu Lys Cys Cys Ser  
20

<210> 476  
<211> 22  
<212> PRT  
<213> Conus aurisiacus

<400> 476  
Met Cys Cys Gly Glu Gly Arg Lys Cys Pro Ser Tyr Phe Arg Asn Ser  
1 5 10 15

Gln Ile Cys His Cys Cys  
20

<210> 477  
<211> 19  
<212> PRT  
<213> Conus aurisiacus

<400> 477  
Cys Cys Arg Trp Pro Cys Pro Arg Gln Ile Asp Gly Glu Tyr Cys Gly  
1 5 10 15

Cys Cys Leu

<210> 478  
<211> 22  
<212> PRT  
<213> Conus bullatus

<400> 478  
Arg Cys Cys Gly Glu Gly Leu Thr Cys Pro Arg Tyr Trp Lys Asn Ser  
1 5 10 15

Gln Ile Cys Ala Cys Cys  
20

<210> 479  
<211> 21  
<212> PRT  
<213> Conus characteristicus

<400> 479

124

Cys Cys Gly Pro Gly Gly Ser Cys Pro Val Tyr Phe Arg Asp Asn Phe  
1 5 10 15

Ile Cys Gly Cys Cys  
20

<210> 480  
<211> 23  
<212> PRT  
<213> Conus circumcisus

<400> 480  
Arg Lys Cys Cys Gly Lys Asp Gly Pro Cys Pro Lys Tyr Phe Lys Asp  
1 5 10 15

Asn Phe Ile Cys Gly Cys Cys  
20

<210> 481  
<211> 20  
<212> PRT  
<213> Conus ermineus

<400> 481  
Cys Cys Ser Trp Pro Cys Pro Arg Tyr Ser Asn Gly Lys Leu Val Cys  
1 5 10 15

Phe Cys Cys Leu  
20

<210> 482  
<211> 21  
<212> PRT  
<213> Conus magus

<400> 482  
Cys Cys Gly Pro Gly Gly Ser Cys Pro Val Tyr Phe Arg Asp Asn Phe  
1 5 10 15

Ile Cys Gly Cys Cys  
20

<210> 483  
<211> 22  
<212> PRT  
<213> Conus magus

<400> 483  
Met Cys Cys Gly Glu Ser Ala Pro Cys Pro Ser Tyr Phe Arg Asn Ser  
1 5 10 15

Gln Ile Cys His Cys Cys  
20

<210> 484  
<211> 22  
<212> PRT  
<213> Conus magus

<400> 484  
Glx Lys Cys Cys Gly Pro Gly Gly Ser Cys Pro Val Tyr Phe Thr Asp  
1 5 10 15

Asn Phe Ile Cys Gly Cys  
20

<210> 485  
<211> 23  
<212> PRT  
<213> Conus magus

<400> 485  
Glx Lys Cys Cys Gly Pro Gly Gly Ser Cys Pro Val Tyr Phe Arg Asp  
1 5 10 15

Asn Phe Ile Cys Gly Cys Cys  
20

<210> 486  
<211> 23  
<212> PRT  
<213> Conus striatus

<400> 486  
Glx Lys Cys Cys Gly Glu Gly Ser Ser Cys Pro Lys Tyr Phe Lys Asn  
1 5 10 15

Asn Phe Ile Cys Gly Cys Cys  
20

<210> 487  
<211> 22  
<212> PRT  
<213> Conus magus

<400> 487  
Glx Lys Cys Cys Ser Gly Gly Ser Cys Pro Leu Tyr Phe Arg Asp Arg  
1 5 10 15

Leu Ile Cys Pro Cys Cys  
20

<210> 488  
<211> 23  
<212> PRT  
<213> Conus stercusmuscarum

<400> 488  
Glx Lys Cys Cys Gly Pro Gly Ala Ser Cys Pro Arg Tyr Phe Lys Asp  
1 5 10 15

Asn Phe Ile Cys Gly Cys Cys  
20

<210> 489  
<211> 22  
<212> PRT  
<213> Conus consors

<400> 489  
Met Cys Cys Gly Glu Gly Ala Pro Cys Pro Ser Tyr Phe Arg Asn Ser  
1 5 10 15

Gln Ile Cys His Cys Cys  
20

<210> 490  
<211> 23  
<212> PRT  
<213> Conus aurisiacus

<400> 490  
Glx Lys Cys Cys Thr Gly Lys Lys Gly Ser Cys Ser Gly Lys Ala Cys  
1 5 10 15

Lys Asn Leu Lys Cys Cys Ser  
20

<210> 491  
<211> 23  
<212> PRT  
<213> Conus aurisiacus

<400> 491  
Glx Lys Cys Cys Thr Gly Arg Lys Gly Ser Cys Ser Gly Lys Ala Cys  
1 5 10 15

Lys Asn Leu Lys Cys Cys Ser  
20

<210> 492  
<211> 23  
<212> PRT  
<213> Conus bullatus

<400> 492  
Val Thr Asp Arg Cys Cys Lys Gly Lys Arg Glu Cys Gly Arg Trp Cys  
1 5 10 15

Arg Asp His Ser Arg Cys Cys  
20

<210> 493  
<211> 23  
<212> PRT  
<213> Conus bullatus

<400> 493  
Val Gly Asp Arg Cys Cys Lys Gly Lys Arg Gly Cys Gly Arg Trp Cys  
1 5 10 15

Arg Asp His Ser Arg Cys Cys  
20

<210> 494  
<211> 24  
<212> PRT  
<213> Conus bullatus

<400> 494  
Val Gly Glu Arg Cys Cys Lys Asn Gly Lys Arg Gly Cys Gly Arg Trp  
1 5 10 15

Cys Arg Asp His Ser Arg Cys Cys  
20

<210> 495  
<211> 26  
<212> PRT  
<213> Conus bullatus

<400> 495  
Ile Val Asp Arg Cys Cys Asn Lys Gly Asn Gly Lys Arg Gly Cys Ser  
1 5 10 15

Arg Trp Cys Arg Asp His Ser Arg Cys Cys

127

20

25

<210> 496  
<211> 25  
<212> PRT  
<213> *Conus bullatus*

<400> 496  
Val Gly Cys Cys Arg Pro Lys Pro Asn Gly Gln Met Met Cys Asp Arg  
1 5 10 15

Trp Cys Glu Lys Asn Ser Arg Cys Cys  
20 25

<210> 497  
<211> 22  
<212> PRT  
<213> *Conus characteristicus*

<400> 497  
Arg Asp Cys Cys Thr Pro Pro Lys Lys Cys Lys Asp Arg Gln Cys Lys  
1 5 10 15

Pro Gln Arg Cys Cys Ala  
20

<210> 498  
<211> 23  
<212> PRT  
<213> *Conus lynceus*

<400> 498  
Gly Arg Asp Cys Cys Thr Pro Pro Arg Lys Cys Arg Asp Arg Ala Cys  
1 5 10 15

Lys Pro Gln Arg Cys Cys Gly  
20

<210> 499  
<211> 22  
<212> PRT  
<213> *Conus lynceus*

<400> 499  
Glx Arg Leu Cys Cys Gly Phe Pro Lys Ser Cys Arg Ser Arg Gln Cys  
1 5 10 15

Lys Pro His Arg Cys Cys  
20

<210> 500  
<211> 22  
<212> PRT  
<213> *Conus laterculatus*

<400> 500  
Arg Asp Cys Cys Thr Pro Pro Lys Lys Cys Arg Asp Arg Gln Cys Lys  
1 5 10 15

Pro Ala Arg Cys Cys Gly  
20

<210> 501  
<211> 22  
<212> PRT

<213> Conus laterculatus

<400> 501

Arg Pro Pro Cys Cys Thr Tyr Asp Gly Ser Cys Leu Lys Glu Ser Cys  
1 5 10 15

Met Arg Lys Ala Cys Cys  
20

<210> 502

<211> 22

<212> PRT

<213> Conus laterculatus

<400> 502

Arg Pro Pro Cys Cys Thr Tyr Asp Gly Ser Cys Leu Lys Glu Ser Cys  
1 5 10 15

Lys Arg Lys Ala Cys Cys  
20

<210> 503

<211> 22

<212> PRT

<213> Conus geographus

<220>

<221> PEPTIDE

<222> (1)..(22)

<223> Xaa is Hyp

<400> 503

Arg Asp Cys Cys Thr Xaa Xaa Lys Lys Cys Lys Asp Arg Gln Cys Lys  
1 5 10 15

Xaa Gln Arg Cys Cys Ala  
20

<210> 504

<211> 22

<212> PRT

<213> Conus geographus

<220>

<221> PEPTIDE

<222> (1)..(22)

<223> Xaa is Hyp

<400> 504

Arg Asp Cys Cys Thr Xaa Xaa Arg Lys Cys Lys Asp Arg Arg Cys Lys  
1 5 10 15

Xaa Met Lys Cys Cys Ala  
20

<210> 505

<211> 22

<212> PRT

<213> Conus geographus

<220>

<221> PEPTIDE

<222> (1)..(22)

<223> Xaa is Hyp

<400> 505  
Arg Asp Cys Cys Thr Xaa Xaa Lys Lys Cys Lys Asp Arg Arg Cys Lys  
1 5 10 15

Xaa Leu Lys Cys Cys Ala  
20

<210> 506  
<211> 22  
<212> PRT  
<213> Conus purpurascens

<220>  
<221> PEPTIDE  
<222> (1)..(22)  
<223> Xaa is Hyp

<400> 506  
Glx Arg Leu Cys Cys Gly Phe Xaa Lys Ser Cys Arg Ser Arg Gln Cys  
1 5 10 15

Lys Xaa His Arg Cys Cys  
20

<210> 507  
<211> 22  
<212> PRT  
<213> Conus magus

<400> 507  
Arg Asp Cys Cys Thr Pro Pro Lys Lys Cys Lys Asp Arg Gln Cys Lys  
1 5 10 15

Pro Gln Arg Cys Cys Ala  
20

<210> 508  
<211> 24  
<212> PRT  
<213> Conus marmoreus

<400> 508  
Arg Gly Gly Cys Cys Thr Pro Pro Arg Lys Cys Lys Asp Arg Ala Cys  
1 5 10 15

Lys Pro Ala Arg Cys Cys Gly Pro  
20

<210> 509  
<211> 23  
<212> PRT  
<213> Conus nobilis

<400> 509  
Glx Lys Cys Cys Thr Gly Lys Lys Gly Ser Cys Ser Gly Lys Ala Cys  
1 5 10 15

Lys Asn Leu Lys Cys Cys Ser  
20

<210> 510  
<211> 24  
<212> PRT  
<213> Conus parius

130

<400> 510  
Arg Gly Gly Cys Cys Thr Pro Pro Lys Lys Cys Lys Asp Arg Ala Cys  
1 5 10 15

Lys Pro Ala Arg Cys Cys Gly Pro  
20

<210> 511  
<211> 23  
<212> PRT  
<213> Conus parius

<400> 511  
Arg Gly Cys Cys Thr Pro Pro Arg Lys Cys Lys Asp Arg Ala Cys Lys  
1 5 10 15

Pro Ala Arg Cys Cys Gly Pro  
20

<210> 512  
<211> 24  
<212> PRT  
<213> Conus radiatus

<220>  
<221> PEPTIDE  
<222> (1)..(24)  
<223> Xaa is Hyp

<400> 512  
Leu Xaa Ser Cys Cys Ser Leu Asn Leu Arg Leu Cys Xaa Val Xaa Ala  
1 5 10 15

Cys Lys Arg Asn Xaa Cys Cys Thr  
20

<210> 513  
<211> 24  
<212> PRT  
<213> Conus radiatus

<220>  
<221> PEPTIDE  
<222> (1)..(24)  
<223> Xaa is Hyp

<400> 513  
Glx Gln Arg Cys Cys Thr Val Lys Arg Ile Cys Xaa Val Xaa Ala Cys  
1 5 10 15

Arg Ser Lys Xaa Cys Cys Lys Ser  
20

<210> 514  
<211> 24  
<212> PRT  
<213> Conus radiatus

<400> 514  
Arg Gly Gly Cys Cys Thr Pro Pro Arg Lys Cys Lys Asp Arg Ala Cys  
1 5 10 15

Lys Pro Ala Arg Cys Cys Gly Pro  
20

<210> 515  
<211> 23  
<212> PRT  
<213> *Conus stercusmuscarum*

<400> 515  
Glx Lys Cys Cys Thr Gly Lys Lys Gly Ser Cys Ser Gly Lys Ala Cys  
1 5 10 15  
Lys Asn Leu Lys Cys Cys Ser  
20

<210> 516  
<211> 21  
<212> PRT  
<213> *Conus tulipa*

<220>  
<221> PEPTIDE  
<222> (1)..(21)  
<223> Xaa is Hyp  
  
<400> 516  
His Gly Cys Cys Lys Gly Xaa Glu Gly Cys Ser Ser Arg Glu Cys Arg  
1 5 10 15  
Xaa Gln His Cys Cys  
20

<210> 517  
<211> 21  
<212> PRT  
<213> *Conus tulipa*

<400> 517  
His Gly Cys Cys Glu Gly Pro Lys Gly Cys Ser Ser Arg Glu Cys Arg  
1 5 10 15  
Pro Gln His Cys Cys  
20

<210> 518  
<211> 23  
<212> PRT  
<213> *Conus wittigi*

<400> 518  
Leu Pro Ser Cys Cys Asp Phe Glu Arg Leu Cys Val Val Pro Ala Cys  
1 5 10 15

Ile Arg His Gln Cys Cys Thr  
20

<210> 519  
<211> 17  
<212> PRT  
<213> *Conus omaria*

<400> 519  
Cys Cys Lys Tyr Gly Trp Thr Cys Leu Leu Gly Cys Thr Pro Cys Asp  
1 5 10 15

Cys

<210> 520

132

<211> 17  
<212> PRT  
<213> Conus omaria

<400> 520  
Cys Cys Arg Tyr Gly Trp Thr Cys Trp Leu Gly Cys Thr Pro Cys Gly  
1 5 10 15

Cys